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(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization
International Bureau



(43) International Publication Date
28 March 2002 (28.03.2002)

PCT

(10) International Publication Number
WO 02/24867 A2

(51) International Patent Classification⁷:**C12N**

(21) International Application Number: PCT/US01/29798

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(22) International Filing Date:

24 September 2001 (24.09.2001)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

09/668,644	22 September 2000 (22.09.2000)	US
09/905,390	13 July 2001 (13.07.2001)	US
09/905,491	13 July 2001 (13.07.2001)	US

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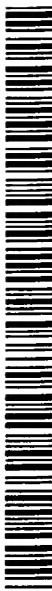
(81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.

(84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

— without international search report and to be republished upon receipt of that report

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

**WO 02/24867 A2**

(54) Title: NOVEL COMPOSITIONS AND METHODS FOR LYMPHOMA AND LEUKEMIA

(57) Abstract: The present invention relates to novel sequences for use in diagnosis and treatment of lymphoma and leukemia. In addition, the present invention describes the use of novel compositions for use in screening methods.

NOVEL COMPOSITIONS AND METHODS FOR LYMPHOMA AND LEUKEMIA

This application is a continuing application of U.S. Serial Number 09/668,644, filed September 22, 2000; U.S. Serial No. 09/905,390, filed July 13, 2001; U.S. Serial No. 09/905,491, filed July 13, 2001; Methods for Diagnosis and Treatment of Diseases Associated with Altered Expression of Pik3r1, filed September 24, 2001; Methods for Diagnosis and Treatment of Diseases Associated with Altered Expression of JAK1, filed September 24, 2001; Methods for Diagnosis and Treatment of Diseases Associated with Altered Expression of Neurogranin, filed September 24, 2001; Methods for Diagnosis and Treatment of Diseases Associated with Altered Expression of Nrf2, filed September 24, 2001; all of which are expressly incorporated herein by reference.

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FIELD OF THE INVENTION

The present invention relates to novel sequences for use in diagnosis and treatment of lymphoma and leukemia, as well as the use of the novel compositions in screening methods.

BACKGROUND OF THE INVENTION

Lymphomas are a collection of cancers involving the lymphatic system and are generally categorized as Hodgkin's disease and Non-Hodgkin lymphoma. Hodgkin's lymphomas are of B lymphocyte origin. Non-Hodgkin lymphomas are a collection of over 30 different types of cancers including T and B lymphomas. Leukemia is a disease of the blood forming tissues and includes B and T cell lymphocytic leukemias. It is characterized by an abnormal and persistent increase in the number of leukocytes and the amount of bone marrow, with enlargement of the spleen and lymph nodes.

Oncogenes are genes that can cause cancer. Carcinogenesis can occur by a wide variety of mechanisms, including infection of cells by viruses containing oncogenes, activation of protooncogenes in the host genome, and mutations of protooncogenes and tumor suppressor genes.

There are a number of viruses known to be involved in human cancer as well

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as in animal cancer. Of particular interest here are viruses that do not contain oncogenes themselves; these are slow-transforming retroviruses. They induce tumors by integrating into the host genome and affecting neighboring protooncogenes in a variety of ways, including promoter insertion, enhancer insertion, and/or truncation of a protooncogene or tumor suppressor gene. The analysis of sequences at or near the insertion sites led to the identification of a number of new protooncogenes.

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With respect to lymphoma and leukemia, murine leukemia retrovirus (MuLV), such as SL3-3 or Akv, is a potent inducer of tumors when inoculated into susceptible newborn mice, or when carried in the germline. A number of sequences have been identified as relevant in the induction of lymphoma and leukemia by analyzing the insertion sites; see Sorensen et al., J. of Virology 74:2161 (2000); Hansen et al., Genome Res. 10(2):237-43 (2000); Sorensen et al., J. Virology 70:4063 (1996); Sorensen et al., J. Virology 67:7118 (1993); Joosten et al., Virology 268:308 (2000); and Li et al., Nature Genetics 23:348 (1999); all of which are expressly incorporated by reference herein.

Accordingly, it is an object of the invention to provide sequences involved in oncogenesis, particularly with respect to lymphomas.

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In this regard, the present invention provides a mammalian Pik3r1 gene which is shown herein to be involved in lymphoma.

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The phosphatidyl inositol 3'-kinases (PI3K, PI3 kinase) represent a ubiquitous family of heterodimeric lipid kinases that are found in association with the cytoplasmic domain of hormone and growth factor receptors and oncogene products. PI3Ks act as downstream effectors of these receptors, are recruited upon receptor stimulation and mediate the activation of second messenger signaling pathways through the production of phosphorylated derivatives of inositol (reviewed in Fry, Biochim. Biophys. Acta., 1226:237-268, 1994). There are multiple forms of PI3K having distinct mechanisms of regulation and different substrate specificities (reviewed in Carpenter et al., Curr. Opin. Biol. 8:153-158, 1996; Zvelebil et al., Phil. Trans. R. Soc. Lond. 351:217-223, 1996).

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The PI3K heterodimers consist of a 110kD (p110) catalytic subunit associated with an 85 kD (Pik3r1) regulatory subunit, and it is through the SH2 domains of the p85 regulatory subunit that the enzyme associates with membrane-bound receptors (Escobedo et al., Cell 65:75-82, 1991; Skolnik et al., Cell 65:83-90, 1991).

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Pik3r1 was originally isolated from bovine brain and shown to exist in two forms, α and β . In these studies, p85 isoforms were shown to bind to and act as substrates for tyrosine-phosphorylated receptor kinases and the polyoma virus middle T antigen complex (Otsu et al., Cell 65:910104, 1991). Since then, the Pik3r1 subunit has been further characterized and shown to interact with a diverse group of proteins including receptor tyrosine kinases such as the erythropoietin receptor, the PDGR- β

receptor and Tie2, an endothelium-specific receptor involved in vascular development and tumor angiogenesis (He et al., Blood 82:3530-3538, 1993; Kontos et al., MCB 18:4131-4140, 1998; Escobedo et al., Cell 65:75-82, 1991). Pik3r1 also interacts with focal adhesion kinase (FAK), a cytoplasmic tyrosine kinase that is involved in integrin signaling, and is thought to be a substrate and effector of FAK. 5
Pik3r1 also interacts with profilin, an actin-binding protein that facilitates actin polymerization (Bhagat et al., Biochem. Mol. Biol. Int. 46:241-248, 1998; Chen et al., PNAS 91:10148-10152, 1994) and the
Pik3r1/profilin complex inhibits actin polymerization.

PI3K has been implicated in the regulation of many cellular activities, including but not limited to survival, proliferation, apoptosis, DNA synthesis, protein transport and neurite extension (reviewed in 10 Fry, supra).

A truncated form of Pik3r1 including the first 571 amino acids of the native protein (as encoded by nucleotides 43-1755 in SEQ ID NO:3 and at Genbank accession number M61906) fused to an amino acid sequence conserved in the eph family of receptor tyrosine kinases causes constitutive activation of PI3K and contributes to cellular transformation of mammalian fibroblasts.

15 A dominant negative isoform of PI3K which inhibits downstream signaling to PKB (Akt) has been isolated (Burgering et al., Nature 376:599-602, 1995). In addition, a constitutively active form of PI3K has been isolated (Klipper et al., MCB 16:4117-4127, 1996; Mante et al., Curr. Biol. 7:63-70, 1996; Franke et al., Cell 81:727-736, 1995).

20 Many approaches to the inhibition of PI3K activity have focussed on the use of inhibitors. Several inhibitors of PI3K activity are known in the literature. These include wortmannin, a fungal metabolite (Ui et al., Trends Biochem. Sci., 20:303-307, 1995), demethoxyviridin, an antifungal agent (Woscholski et al., FEBS Lett. 342:109-114, 1994), quercetin and LY294002 (Vlahos et al., JBC 269:5241-5248, 1994). These inhibitors primarily target the p110 subunit of PI3k.

25 An additional approach taken to inhibit PI3K activity involves the inhibition of Pik3r1 expression, as through the use of antisense oligonucleotides directed to Pik3r1 nucleic acid sequence (for example, see US Patent 6,100,090 issued to Monia et al.).

As disclosed herein, alteration and/or dysregulation of Pik3r1 leads to lymphoma. Provided herein are novel compositions and methods for the diagnosis, treatment, and prophylaxis of lymphoma.

30 As demonstrated herein, GNAS genes are also implicated in lymphomas and leukemias. GNAS is a complex locus encoding multiple proteins, including an α subunit of a stimulatory G protein ($G_s\alpha$). G proteins transduce extracellular signals in signal transduction pathways. Each G protein is a heterotrimer, composed of an α , β and γ subunit. The β and γ subunits anchor the protein to the

cytoplasmic side of the plasma membrane. Upon binding of a ligand, G_sα dissociates from the complex, transducing signals from hormone receptors to effector molecules including adenylyl cyclase resulting in hormone-stimulated cAMP generation (Molecular Biology of the Cell, 3d edition, Alberts, B et al., Garland Publishing 1994).

5 Other proteins generated from the GNAS locus, through alternative splicing, include XLαs, a G_sα isoform with an extended NH₂ terminal extension, and NESP55, a chromogranin-like neurosecretory protein (Weinstein LS et al., Am J Physiol Renal Physiol 2000, 278:F507-14). In mice, Nesp, the mouse homolog of NESP55, is located 15 kb upstream of Gnasxl, the mouse homolog of XLαs, which is in turn, 30 kb upstream of Gnas (Wroe et al., Proc. Natl. Acad. Sci. 97:3342 (2000)). NESP55 is
10 processed into smaller peptides , one of which acts as an inhibitor of the serotonergic 5-HT_{1B} receptor (Ischia et. al. J. Biol. Chem. 272:11657 (1997)). The function of XLαs is not known, but it is also expressed primarily in the neuroendocrine system and may be involved in pseudohypoparathyroidism type Ia (Hayward et al., Proc. Natl. Acad. Sci. 95:10038 (1998)). XLαs and NESP55 have been found to be expressed in opposite parental alleles, as a result of imprinting (Wroe et al., Proc. Natl. Acad.
15 Sci. 97:3342 (2000)).

GNAS also plays a role in diseases other than leukemias and lymphomas. Mutations in GNAS1, the human GNAS gene, result in Albright hereditary osteodystrophy (AHO), a disease characterized by short stature and obesity. Studies with the mouse homolog demonstrate that the obesity seen is a consequence of the reduced expression of GNAS. In contrast, other mutations have been shown to
20 result in constitutive activation of G_sα, resulting in endocrine tumors and McCune-Albright syndrome, a condition characterized by abnormalities in endocrine function (Aldred MA and Trembath, RC, Hum Mutat 2000, 16:183-9). The mechanism behind this disease as well as fibrous dysplasia, a progressive bone disease, is caused by increased cAMP levels which results in increase IL-6 levels, triggering abnormal osteoblast differentiation and increased osteoclastic activity (Stanton RP et al., J. Bone
25 Miner. Res. 1999, 14:1104-14).

Accordingly, it is an object of the invention to provide methods for detection and screening of drug candidates for diseases involving GNAS, particularly with respect to lymphomas.

As demonstrated herein, a HIPK1 gene is also implicated in lymphomas and leukemias. HIPK1 is a member of a novel family of nuclear protein kinases that act as transcriptional co-repressors for NK
30 class of homeoproteins (Kim YH et al., J. Biol. Chem. 1998, 273:25875-25879). Homeoproteins are transcription factors that regulate homeobox genes, which are involved in various developmental processes, such as pattern formation and organogenesis (McGinnis, W. and Krumlauf, R., Cell 1992, 68:283-302).

Homeoproteins may play a role in human disease. Aberrant expression of the NKX2-5 homeodomain transcription factor has been found to be involved in a congenital heart disease (Schott, J.-J. et al., Science 1998, 281:108-111).

5 Accordingly, it is an object of the invention to provide methods for detection and screening of drug candidates for diseases involving HIPK1; particularly with respect to lymphomas.

Cytokines and Interferons regulate a wide range of cellular functions in the lympho-hematopoietic system. This regulation is mediated, in part, by the Jak-STAT pathway. In this pathway a Cytokine or Interferon initially binds to the extracellular portion of a membrane bound receptor. Binding of a Cytokine or Interferon activates members of the Janus family of Tyrosine Kinases (JAKs), including 10 JAK1. Activated JAKs phosphorylate docking sites on the intracellular portion of the receptor which in turn activate transcription factors known as the signal transducers and activators of transcription (STATs). Once activated, STATs dimerize and translocate to the nucleus to bind target DNA sequences resulting in modulation of gene expression.

Given the integral role JAKs play in this signal transduction pathway it is not surprising that a number 15 of studies have shown that JAK dysregulation leads to severe disease states. JAK mutations in Drosophila termed *Tum-I*, Tumorous lethal, for example, lead to leukemia in flies. Harrison et al., EMBO J. 14:1412-20 (1995); Luo et al., EMBO J. 14:1412-20 (1995); Luo et al., Mol. Cell Biol. 17:1562-71 (1997). Additionally, constitutive activation of JAKs in mammalian cells has been shown to lead to malignant transformation in several settings. Migone et al., Science 269:79-81 (1995); 20 Zhang et al., Proc. Natl. Acad. Sci. USA 93:9148-53 (1996); Danial et al., Science 269:1875-77 (1995); Meydan et al., Nature 379:645-48 (1996). Accordingly, understanding the various aspects of JAK function, its binding capabilities, catalytic aspects, etc., will give insight into a number of disease states not the least of which being either lymphoma or leukemia.

25 Neurogranin is a neuronal protein thought to play a role in dendritic spine formation and synaptic plasticity. The Neurogranin gene encodes a 78-amino acid protein that functions as a postsynaptic kinase substrate and has been shown to bind calmodulin in the absence of calcium. Martinez de Arrieta et al., Endocrinology 140(1):335-43 (1999). Though little is understood at the present time, dysregulation of Neurogranin gene expression has been implicated in disease states. Recent studies 30 have shown Neurogranin expression is tightly regulated by thyroid hormone. Morte et al., FEBS Lett Dec 31; 464(3):179-83 (1999). This regulation may explain the role hypothyroidism has on mental states during development as well as in adult subjects. Additionally, a transactivator overexpressed in prostate cancer, EGR1, has been shown to induce Neurogranin which may explain the neuroendocrine differentiation that often accompanies prostate cancer progression. Svaren et al., J. Biol. Chem. Dec 8; 275(49):38524-31 (2000). Accordingly, understanding the various aspects of

Neurogranin structure and function will likely lead to a clearer view of its role in hypothyroidism and prostate cancer, as well as other diseases such as lymphoma and leukemia.

Accordingly, it is an object of the invention to provide compositions involved in oncogenesis, particularly with respect to the role of Neurogranin in lymphomas.

- 5 Also, in this regard, the present invention provides a mammalian Nrf2 gene which is shown herein to be involved in lymphoma.

The Nrf2 gene encodes a DNA binding transcriptional regulatory protein (transcription factor) belonging to the "cap 'n collar" subfamily of the basic leucine zipper family of transcription factors (Chan et al., PNAS 93:13943-13948, 1996; Moi et al., PNAS 91:9926-9930, 1994). The Nrf2 gene produces a 2.2kb transcript which predicts a 66 kDa protein (Moi et al., PNAS 91:9926-9930, 1994).
10 The Nrf2 protein binds to a DNase hypersensitive site located in the β-globin locus control region (Moi et al., PNAS 91:9926-9930, 1994), as well as to the antioxidant response element (ARE) which is found in the regulatory regions of many detoxifying enzyme genes (Venugopal et al., Oncogene, 17:3145-3156, 1998).

15 Nrf2 gene function is not required for normal development, as evidenced by homozygous disruption of the Nrf2 loci in transgenic mice (Chan et al., PNAS 93:13943-13948, 1996). However, loss of Nrf2 gene function compromises the ability of haematopoietic cells to endure oxidative stress (Ishii et al., J. Biol. Chem., 275:16023-16029, 2000; Enomoto et al., Toxicol. Sci., 59:169-177, 2001) and sensitizes cells to the carcinogenic activity of oxidative agents (Ramos-Gomez et al., PNAS, 98:3410-3415,
20 2001).

Nrf2 proteins are capable of interacting with other transcription factors, including Jun proteins (Venugopal et al., Oncogene, 17:3145-3156, 1998) and Maf proteins (Marini et al., J. Biol. Chem., 272:16490-16497, 1997). Jun proteins appear to cooperate with Nrf2 to regulate the transcription of target genes (Venugopal et al., Oncogene, 17:3145-3156, 1998) while Maf proteins appear to antagonize the transcription promoting activity of Nrf2 protein (Nguyen et al., J. Biol. Chem., 275:15466-15473, 2000). In addition, the human cytomegalovirus protein IE-2 has also been found to interact with Nrf2 and to inhibit its transcription promoting activity (Huang et al., J. Biol. Chem., 275:12313-12320, 2000).

25 Despite being dispensable for the normal development of lymphoid cells and tissues, which includes the normal processes of B cell and T cell determination, differentiation, proliferation, and death, it is demonstrated herein that dysregulation of the Nrf2 gene leads to lymphoma.
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SUMMARY OF THE INVENTION

In accordance with the objects outlined above, the present invention provides methods for screening for compositions which modulate lymphomas. Also provided herein are methods of inhibiting proliferation of a cell, preferably a lymphoma cell. Methods of treatment of lymphomas, including diagnosis, are also provided herein.

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In one aspect, a method of screening drug candidates comprises providing a cell that expresses a lymphoma associated (LA) gene or fragments thereof. Preferred embodiments of LA genes are genes which are differentially expressed in cancer cells, preferably lymphoma or leukemia cells, compared to other cells. Preferred embodiments of LA genes used in the methods herein include, but 10 are not limited to the nucleic acids selected from Tables 1, 2 or 3. Additional preferred embodiments include, but are not limited to, the nucleic acids set forth in Tables 4, 6, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 22, 23, 24, 27, 28 or 30. The method further includes adding a drug candidate to the cell and determining the effect of the drug candidate on the expression of the LA gene.

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In one embodiment, the method of screening drug candidates includes comparing the level of expression in the absence of the drug candidate to the level of expression in the presence of the drug candidate.

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Also provided herein is a method of screening for a bioactive agent capable of binding to a LA protein (LAP), the method comprising combining the LAP and a candidate bioactive agent, and determining the binding of the candidate agent to the LAP. In a preferred embodiment, a LA protein is selected from the amino acid sequences set forth in Tables 5, 7, 9, 10, 11, 12, 13, 14, 16, 17, 20, 21, 25, 26, 29 or 31.

Further provided herein is a method for screening for a bioactive agent capable of modulating the activity of a LAP. In one embodiment, the method comprises combining the LAP and a candidate bioactive agent, and determining the effect of the candidate agent on the bioactivity of the LAP.

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Also provided is a method of evaluating the effect of a candidate lymphoma drug comprising administering the drug to a patient and removing a cell sample from the patient. The expression profile of the cell is then determined. This method may further comprise comparing the expression profile of the patient to an expression profile of a healthy individual.

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In a further aspect, a method for inhibiting the activity of an LA protein is provided. In one embodiment, the method comprises administering to a patient an inhibitor of an LA protein preferably encoded by a nucleic acid selected from the group consisting of the sequences outlined in Tables 1, 2 or 3. Additional preferred embodiments include, but are not limited to, the nucleic acids set forth in Tables 4, 6, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 22, 23, 24, 27, 28 or 30. In a preferred

embodiment, a LA protein is selected from the amino acid sequences set forth in Tables 5, 7, 9, 10, 11, 12, 13, 14, 16, 17, 20, 21, 25, 26, 29 or 31.

5 A method of neutralizing the effect of a LA protein, preferably selected from the group of sequences outlined in Tables, 1, 2 or 3, is also provided. Additional preferred embodiments include, but are not limited to, the nucleic acids set forth in Tables 4, 6, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 22, 23, 24, 27, 28 or 30. In a preferred embodiment, a LA protein is selected from the amino acid sequences set forth in Tables 5, 7, 9, 10, 11, 12, 13, 14, 16, 17, 20, 21, 25, 26, 29 or 31. Preferably, the method comprises contacting an agent specific for said protein with said protein in an amount sufficient to effect neutralization.

10 Moreover, provided herein is a biochip comprising a nucleic acid segment which encodes a LA protein, preferably selected from the sequences outlined in Tables 1, 2 or 3. Additional preferred embodiments include, but are not limited to, the nucleic acids set forth in Tables 4, 6, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 22, 23, 24, 27, 28 or 30. In a preferred embodiment, a LA protein is selected from the amino acid sequences set forth in Tables 5, 7, 9, 10, 11, 12, 13, 14, 16, 17, 20, 21, 25, 26, 29 or 31.

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Also provided herein is a method for diagnosing or determining the propensity to lymphomas by sequencing at least one LA gene of an individual. In yet another aspect of the invention, a method is provided for determining LA gene copy number in an individual.

20 Novel sequences are also provided herein. Other aspects of the invention will become apparent to the skilled artisan by the following description of the invention.

In one aspect the present invention provides an LA protein known as Pik3r1 comprising the amino acid sequence set forth in SEQ ID NO:179 and at Genbank Accession number AAC52847, which is encoded by the Pik3r1 nucleic acid sequence set forth by nucleotides 575 to 2749 in SEQ ID NO:178 and at Genbank Accession Number U50413. In one aspect the present invention provides an LA nucleic acid referred to herein as Pik3r1 and comprising the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank Accession number U50413, which encodes an Pik3r1 protein.

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In one aspect the present invention provides an LA protein known as Pik3r1 comprising the amino acid sequence set forth in SEQ ID NO:181 and at Genbank Accession number A38748. In one aspect the present invention provides an LA nucleic acid referred to herein as Pik3r1 and comprising the nucleic acid sequence set forth by nucleotides 43 to 2217 in SEQ ID NO:3 and at Genbank Accession number M61906, which encodes an Pik3r1 protein.

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Also provided herein are Pik3r1 nucleic acids comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank Accession number U50413, or complements thereof.

Also provided herein are Pik3r1 nucleic acids comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank accession number M61906, or complements thereof.

5 Also provided herein are Pik3r1 nucleic acids which will hybridize under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank accession number U50413, or complements thereof.

Also provided herein are Pik3r1 nucleic acids which will hybridize under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank accession number M61906, or complements thereof.

10 Also provided herein are Pik3r1 proteins encoded by Pik3r1 nucleic acids as described herein.

Also provided herein are Pik3r1 proteins comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:179 and at Genbank accession number AAC52847.

15 Also provided herein are Pik3r1 proteins comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:181 and at Genbank accession number A38748.

Also provided herein are Pik3r1 genes encoding Pik3r1 proteins comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:179 and at Genbank accession number AAC52847.

20 Also provided herein are Pik3r1 genes encoding Pik3r1 proteins comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:181 and at Genbank accession number A38748.

In one aspect, the present invention provides a method for screening for a candidate bioactive agent capable of modulating the activity of a Pik3r1 gene. In one embodiment, such a method comprises adding a candidate agent to a cell and determining the level of expression of a Pik3r1 gene in the presence and absence of the candidate agent. In a preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank accession number U50413. In another preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank accession number M61906.

30 Further provided herein is a method for screening for a candidate bioactive agent capable of modulating the activity of a Pik3r1 protein encoded by a Pik3r1 gene. In one embodiment, such a method comprises contacting a Pik3r1 protein or a cell comprising a Pik3r1 protein, and a candidate

bioactive agent, and determining the effect on the activity of the Pik3r1 protein in the presence and absence of the candidate agent. In another embodiment, such a method comprises contacting a cell comprising a Pik3r1 protein, and a candidate bioactive agent, and determining the effect on the cell in the presence and absence of the candidate agent. In a preferred embodiment, a Pik3r1 protein 5 comprises the amino acid sequence set forth in SEQ ID NO:179 and at Genbank accession number AAC52847, or a fragment thereof. In another preferred embodiment, a Pik3r1 protein comprises the amino acid sequence set forth in SEQ ID NO:181 and at Genbank accession number A38748, or a fragment thereof. In a preferred embodiment, a Pik3r1 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank accession number 10 U50413, or a fragment thereof. In another preferred embodiment, a Pik3r1 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank accession number M61906, or a fragment thereof. In one embodiment, a Pik3r1 protein is a recombinant protein. In one embodiment, a Pik3r1 protein is isolated. In one embodiment, a Pik3r1 protein is cell-free, as in a cell lysate.

15 Also provided herein is a method for screening for a bioactive agent capable of binding to a Pik3r1 protein encoded by a Pik3r1 gene. In one embodiment, such a method comprises combining a Pik3r1 protein or a cell comprising a Pik3r1 protein, and a candidate bioactive agent, and determining the binding of the candidate agent to the Pik3r1 protein. In a preferred embodiment, a Pik3r1 protein comprises the amino acid sequence set forth in SEQ ID NO:179, or a fragment thereof. In another 20 preferred embodiment, a Pik3r1 protein comprises the amino acid sequence set forth in SEQ ID NO:181, or a fragment thereof. In a preferred embodiment, a Pik3r1 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:178, or a fragment thereof. In another preferred embodiment, a Pik3r1 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:180, or a fragment thereof. In one embodiment, a 25 Pik3r1 protein is a recombinant protein. In one embodiment, a Pik3r1 protein is isolated. In one embodiment, a Pik3r1 protein is cell-free, as in a cell lysate.

Also provided is a method for evaluating the effect of a candidate lymphoma drug, comprising administering the drug to a patient and removing a cell sample or a cell fraction sample from the patient. A gene expression profile for the sample is then determined, including determination of the 30 expression of a Pik3r1 gene. In a preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence set forth in SEQ ID NO:178, or a fragment thereof. In another preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence set forth in SEQ ID NO:180, or a fragment thereof. Such a method may further comprise comparing the expression profile of the patient sample to an expression profile of a healthy individual sample.

35 In a further aspect, a method for inhibiting the activity of a Pik3r1 protein is provided. In one embodiment, the method comprises administering to a patient an inhibitor of a Pik3r1 protein. In a preferred embodiment, a Pik3r1 protein comprises the amino acid sequence set forth in SEQ ID NO:179 or a fragment thereof. In another preferred embodiment, a Pik3r1 protein comprises the

amino acid sequence set forth in SEQ ID NO:181 or a fragment thereof. In a preferred embodiment, a Pik3r1 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:178 or a fragment thereof. In another preferred embodiment, a Pik3r1 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:180 or a fragment thereof.

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Also provided herein is a method for neutralizing Pik3r1 protein activity with a bioactive agent. In a preferred embodiment, a Pik3r1 protein comprises the amino acid sequence set forth in SEQ ID NO:179 or a fragment thereof. In another preferred embodiment, a Pik3r1 protein comprises the amino acid sequence set forth in SEQ ID NO:181 or a fragment thereof. In a preferred embodiment, a Pik3r1 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:178, or a fragment thereof. In another preferred embodiment, a Pik3r1 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:180, or a fragment thereof. In one embodiment, such a method comprises contacting a Pik3r1 protein with an agent that specifically modulates Pik3r1 protein activity, in an amount sufficient to effect neutralization.

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Moreover, provided herein is a biochip comprising a nucleic acid which encodes a Pik3r1 protein or a portion thereof. In a preferred embodiment, a Pik3r1 nucleic acid comprises the nucleic acid sequence set forth in SEQ ID NO:178, or complement thereof, or a fragment thereof or complement of a fragment thereof. In another preferred embodiment, a Pik3r1 nucleic acid comprises the nucleic acid sequence set forth in SEQ ID NO:180, or complement thereof, or a fragment thereof or complement of a fragment thereof.

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Also provided herein is a method for diagnosing or determining a predisposition for lymphomas, comprising sequencing at least one Pik3r1 gene from an individual and determining the nucleic acid sequence of the Pik3r1 gene or a fragment thereof. In a preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence set forth in SEQ ID NO:178, or a fragment thereof. In another preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence set forth in SEQ ID NO:180, or a fragment thereof.

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Similarly provided are methods for determining lymphoma subtype and determining a prognosis for an individual having lymphoma, which comprise sequencing at least one Pik3r1 gene from an individual and determining the nucleic acid sequence of the Pik3r1 gene or a fragment thereof. In a preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence set forth in SEQ ID NO:178, or a fragment thereof. In another preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence set forth in SEQ ID NO:180, or a fragment thereof.

In yet another aspect of the invention, a method is provided for determining the number of copies of a Pik3r1 gene in an individual. In a preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence set forth in SEQ ID NO:178, or complement thereof, or a fragment thereof or complement of a fragment thereof. In a preferred embodiment, a Pik3r1 gene comprises the nucleic acid sequence

set forth in SEQ ID NO:180, or complement thereof, or a fragment thereof or complement of a fragment thereof.

In yet another aspect of the invention, a method is provided for determining the chromosomal location of a *Pik3r1* gene. In a preferred embodiment, a *Pik3r1* gene comprises the nucleic acid sequence set forth in SEQ ID NO:178, or a fragment thereof. In another preferred embodiment, a *Pik3r1* gene comprises the nucleic acid sequence set forth in SEQ ID NO:180, or a fragment thereof. Such a method may be used to determine *Pik3r1* gene rearrangements or translocations. Without being bound by theory, *Pik3r1* gene rearrangement and translocation events appear to be important in the aetiology of lymphoma.

It is an object of this invention that the identification *Pik3r1* genes and recognition of their involvement in lymphoma provide diagnostic agents to distinguish between lymphoma subtypes, and analytical agents for further analysis of mechanisms involved in dysregulated growth and/or survival and/or apoptosis in cells of the hematopoietic system. An additional object of the invention is to provide appropriate and potentially novel targets for therapeutic interventions, particularly with regard to lymphoma, which are identified through the use of the diagnostic and analytical agents provided herein.

Without being bound by theory, it is recognized herein that the involvement of *Pik3r1* genes in the cellular dysregulation underlying lymphoma implicates genes having products which are regulated by the PI3K pathway, preferably by phosphorylation by protein kinase B (PKB; AKT) and/or protein kinase C (PKC), in the cellular dysregulation underlying lymphoma.

Moreover, it is recognized herein that dysregulated growth in the hematopoietic system has been attributed to the inhibition of apoptosis, for example as by the deregulated expression of Bcl-2. Without being bound by theory, the present disclosure provides a new molecular mechanism for lymphoma in which alterations in *Pik3r1* lead to alterations in the activity of PKB and the phosphorylation of proteins involved in survival and cell death, such as the Bcl-2 family member "BAD" (see Datta et al., Cell 91:231-241, 1997; del Peso et al., Science 278:687-689, 1997).

Novel sequences are also provided herein. Other aspects of the invention will become apparent to the skilled artisan by the following description of the invention.

In one aspect, a method of screening drug candidates comprises providing a cell that expresses a GNAS gene or fragments thereof. The method further includes adding a drug candidate to the cell and determining the effect of the drug candidate on the expression of a GNAS gene.

In one embodiment, the method of screening drug candidates includes comparing the level of expression in the absence of the drug candidate to the level of expression in the presence of the drug candidate.

Also provided herein is a method of screening for a bioactive agent capable of binding to a protein encoded by a GNAS gene, e.g. G_sα, the method comprising combining a Gnas protein and a candidate bioactive agent, and determining the binding of the candidate agent to the Gnas protein.

5 Further provided herein is a method for screening for a bioactive agent capable of modulating the activity of a protein encoded by a GNAS gene. In one embodiment, the method comprises combining a Gnas protein and a candidate bioactive agent, and determining the effect of the candidate agent on the bioactivity of a Gnas protein.

10 Also provided is a method of evaluating the effect of a candidate lymphoma drug comprising administering the drug to a patient and removing a cell sample from the patient. The expression profile of the cell is then determined. This method may further comprise comparing the expression profile of the patient to an expression profile of a healthy individual.

In a further aspect, a method for inhibiting the activity of a protein encoded by a GNAS gene is provided. In one embodiment, the method comprises administering to a patient an inhibitor of a Gnas protein.

15 A method of neutralizing the effect of Gnas proteins is also provided. Preferably, the method comprises contacting an agent specific for said protein with said protein in an amount sufficient to effect neutralization.

Moreover, provided herein is a biochip comprising a nucleic acid segment which encodes a Gnas protein.

20 Also provided herein is a method for diagnosing or determining the propensity to diseases, including lymphomas, by sequencing at least one GNAS gene of an individual. In yet another aspect of the invention, a method is provided for determining GNAS gene copy number in an individual.

25 In one aspect, a method of screening drug candidates comprises providing a cell that expresses a HIPK1 gene or fragments thereof. The method further includes adding a drug candidate to the cell and determining the effect of the drug candidate on the expression of a HIPK1 gene.

In one embodiment, the method of screening drug candidates includes comparing the level of expression in the absence of the drug candidate to the level of expression in the presence of the drug candidate.

30 Also provided herein is a method of screening for a bioactive agent capable of binding to a protein encoded by a HIPK1 gene, the method comprising combining a HIPK1 protein and a candidate bioactive agent, and determining the binding of the candidate agent to a HIPK1 protein.

Further provided herein is a method for screening for a bioactive agent capable of modulating the activity of a protein encoded by a HIPK1 gene. In one embodiment, the method comprises combining a HIPK1 protein and a candidate bioactive agent, and determining the effect of the candidate agent on the bioactivity of a HIPK1 protein.

5 Also provided is a method of evaluating the effect of a candidate lymphoma drug comprising administering the drug to a patient and removing a cell sample from the patient. The expression profile of the cell is then determined. This method may further comprise comparing the expression profile of the patient to an expression profile of a healthy individual.

10 In a further aspect, a method for inhibiting the activity of a protein encoded by a HIPK1 gene is provided. In one embodiment, the method comprises administering to a patient an inhibitor of a HIPK1 protein.

A method of neutralizing the effect of HIPK1 protein is also provided. Preferably, the method comprises contacting an agent specific for said protein with said protein in an amount sufficient to effect neutralization.

15 Moreover, provided herein is a biochip comprising a nucleic acid segment which encodes HIPK1 protein.

Also provided herein is a method for diagnosing or determining the propensity to diseases, including lymphomas, by sequencing at least one HIPK1 gene of an individual. In yet another aspect of the invention, a method is provided for determining HIPK1 gene copy number in an individual.

20 In one aspect, a method of screening drug candidates comprises providing a cell that expresses a JAK1 gene or fragments thereof. Preferred embodiments of JAK1 genes are genes which are differentially expressed in cancer cells, preferably lymphoma or leukemia cells, compared to other cells. The method further includes adding a drug candidate to the cell and determining the effect of the drug candidate on the expression of the JAK1 gene.

25 In one embodiment, the method of screening drug candidates includes comparing the level of expression in the absence of the drug candidate to the level of expression in the presence of the drug candidate.

30 Also provided herein is a method of screening for a bioactive agent capable of binding to a JAK1 protein, the method comprising combining the JAK1 protein and a candidate bioactive agent, and determining the binding of the candidate agent to the JAK1 protein.

Further provided herein is a method for screening for a bioactive agent capable of modulating the activity of JAK1 protein. In one embodiment, the method comprises combining the JAK1 protein and a

candidate bioactive agent, and determining the effect of the candidate agent on the bioactivity of the JAKI protein.

Also provided is a method of evaluating the effect of a candidate lymphoma drug comprising administering the drug to a patient and removing a cell sample from the patient. The expression profile of the cell is then determined. This method may further comprise comparing the expression profile of the patient to an expression profile of a healthy individual.
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In a further aspect, a method for inhibiting the activity of a JAKI protein is provided.

A method of neutralizing the effect of a JAKI protein, is also provided. Preferably, the method comprises contacting an agent specific for said protein with said protein in an amount sufficient to
10 effect neutralization.

Moreover, provided herein is a biochip comprising a nucleic acid segment which encodes a JAKI protein.

Also provided herein is a method for diagnosing or determining the propensity to lymphomas by sequencing the JAKI gene of an individual. In yet another aspect of the invention, a method is
15 provided for determining JAKI gene copy number in an individual.

In one aspect, a method of screening drug candidates comprises providing a cell that expresses a Neurogranin gene or fragments thereof. Preferred embodiments of Neurogranin genes are genes which are differentially expressed in cancer cells, preferably lymphoma or leukemia cells, compared to other cells. The method further includes adding a drug candidate to the cell and determining the effect
20 of the drug candidate on the expression of the Neurogranin gene.

In one embodiment, the method of screening drug candidates includes comparing the level of expression in the absence of the drug candidate to the level of expression in the presence of the drug candidate.

Also provided herein is a method of screening for a bioactive agent capable of binding to a
25 Neurogranin protein, the method comprising combining the Neurogranin protein and a candidate bioactive agent, and determining the binding of the candidate agent to the Neurogranin protein.

Further provided herein is a method for screening for a bioactive agent capable of modulating the activity of Neurogranin protein. In one embodiment, the method comprises combining the Neurogranin protein and a candidate bioactive agent, and determining the effect of the candidate agent on the bioactivity of the Neurogranin protein.
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Also provided is a method of evaluating the effect of a candidate lymphoma drug comprising administering the drug to a patient and removing a cell sample from the patient. The expression profile of the cell is then determined. This method may further comprise comparing the expression profile of the patient to an expression profile of a healthy individual.

- 5 In a further aspect, a method for inhibiting the activity of a Neurogranin protein is provided. In one embodiment, the method comprises administering to a patient an inhibitor of a Neurogranin protein.

A method of neutralizing the effect of a Neurogranin protein, is also provided. Preferably, the method comprises contacting an agent specific for said protein with said protein in an amount sufficient to effect neutralization.

- 10 Moreover, provided herein is a biochip comprising a nucleic acid segment which encodes a Neurogranin protein.

Also provided herein is a method for diagnosing or determining the propensity to lymphomas by sequencing the Neurogranin gene of an individual. In yet another aspect of the invention, a method is provided for determining Neurogranin gene copy number in an individual.

- 15 In one aspect the present invention provides an LA protein known as Nrf2. In a preferred embodiment Nrf2 comprises the amino acid sequence set forth in SEQ ID NO:211 and at Genbank Accession number AAA68291, which is encoded by the Nrf2 nucleic acid sequence set forth by nucleotides 298 to 2043 in SEQ ID NO:210 and at Genbank Accession Number U20532. In one aspect the present invention provides an LA nucleic acid referred to herein as Nrf2. In a preferred embodiment the Nrf2 nucleic acid comprises the nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank Accession number U20532, which encodes an Nrf2 protein.

- 20 In one aspect the present invention provides an LA protein known as Nrf2 comprising the amino acid sequence set forth in SEQ ID NO:213 and at Genbank Accession number NP_006155, which is encoded by the Nrf2 nucleic acid sequence set forth by nucleotides 40 to 1809 in SEQ ID NO:212 and at Genbank Accession Number NM_006164. In one aspect the present invention provides an LA nucleic acid referred to herein as Nrf2 and comprising the nucleic acid sequence set forth in SEQ ID NO:212 and at Genbank Accession number NM_006164, which encodes an Nrf2 protein.

- 25 Also provided herein are Nrf2 nucleic acids comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank Accession number U20532, or complements thereof.

30 Also provided herein are Nrf2 nucleic acids comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:212 and at Genbank accession number NM_006164, or complements thereof.

Also provided herein are Nrf2 nucleic acids which will hybridize under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank accession number U20532, or complements thereof.

5 Also provided herein are Nrf2 nucleic acids which will hybridize under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:212 and at Genbank accession number NM_006164, or complements thereof.

Also provided herein are Nrf2 proteins encoded by Nrf2 nucleic acids as described herein.

10 Also provided herein are Nrf2 proteins comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:211 and at Genbank accession number AAA68291.

Also provided herein are Nrf2 proteins comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:213 and at Genbank accession number NP_006155.

15 Also provided herein are Nrf2 genes encoding Nrf2 proteins comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:211 and at Genbank accession number AAA68291.

Also provided herein are Nrf2 genes encoding Nrf2 proteins comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:213 and at Genbank accession number NP_006155.

20 In one aspect, the present invention provides a method for screening for a candidate bioactive agent capable of modulating the activity of an Nrf2 gene. In one embodiment, such a method comprises adding a candidate agent to a cell and determining the level of expression of an Nrf2 gene in the presence and absence of the candidate agent. In a preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank accession number U20532. In another preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:212 and at Genbank accession number NM_006164.

25 Further provided herein is a method for screening for a candidate bioactive agent capable of modulating the activity of an Nrf2 protein encoded by an Nrf2 gene. In one embodiment, such a method comprises contacting an Nrf2 protein or a cell comprising an Nrf2 protein, and a candidate bioactive agent, and determining the effect on the activity of the Nrf2 protein in the presence and absence of the candidate agent. In another embodiment, such a method comprises contacting a cell comprising an Nrf2 protein, and a candidate bioactive agent, and determining the effect on the cell in

the presence and absence of the candidate agent. In a preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth in SEQ ID NO:211 and at Genbank accession number AAA68291, or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth in SEQ ID NO:213 and at Genbank accession number NP_006155, or a fragment thereof. In a preferred embodiment, an Nrf2 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank accession number U20532, or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:212 and at Genbank accession number NM_006164, or a fragment thereof. In one embodiment, an Nrf2 protein is a recombinant protein. In one embodiment, an Nrf2 protein is isolated. In one embodiment, an Nrf2 protein is cell-free, as in a cell lysate.

Also provided herein is a method for screening for a bioactive agent capable of binding to an Nrf2 protein encoded by an Nrf2 gene. In one embodiment, such a method comprises combining an Nrf2 protein or a cell comprising an Nrf2 protein, and a candidate bioactive agent, and determining the binding of the candidate agent to the Nrf2 protein. In a preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth in SEQ ID NO:211, or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth in SEQ ID NO:213, or a fragment thereof. In a preferred embodiment, an Nrf2 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:210, or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:212, or a fragment thereof. In one embodiment, an Nrf2 protein is a recombinant protein. In one embodiment, an Nrf2 protein is isolated. In one embodiment, an Nrf2 protein is cell-free, as in a cell lysate.

Also provided is a method for evaluating the effect of a candidate lymphoma drug, comprising administering the drug to a patient and removing a cell sample or a cell fraction sample from the patient. A gene expression profile for the sample is then determined, including determination of the expression of an Nrf2 gene. In a preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:210, or a fragment thereof. In another preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:212, or a fragment thereof. Such a method may further comprise comparing the expression profile of the patient sample to an expression profile of a healthy individual sample.

In a further aspect, a method for inhibiting the activity of an Nrf2 protein is provided. In one embodiment, the method comprises administering to a patient an inhibitor of an Nrf2 protein. In a preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth in SEQ ID NO:211 or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth in SEQ ID NO:213 or a fragment thereof. In a preferred embodiment, an Nrf2 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:210 or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises

an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:212 or a fragment thereof.

Also provided herein is a method for neutralizing Nrf2 protein activity with a bioactive agent. In a preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth in SEQ ID NO:211 or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth in SEQ ID NO:213 or a fragment thereof. In a preferred embodiment, an Nrf2 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:210, or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises an amino acid sequence encoded by the nucleic acid sequence set forth in SEQ ID NO:212, or a fragment thereof. In one embodiment, such a method comprises contacting an Nrf2 protein with an agent that specifically modulates Nrf2 protein activity, in an amount sufficient to effect neutralization.

Moreover, provided herein is a biochip comprising a nucleic acid which encodes an Nrf2 protein or a portion thereof. In a preferred embodiment, an Nrf2 nucleic acid comprises the nucleic acid sequence set forth in SEQ ID NO:210, or complement thereof, or a fragment thereof or complement of a fragment thereof. In another preferred embodiment, an Nrf2 nucleic acid comprises the nucleic acid sequence set forth in SEQ ID NO:212, or complement thereof, or a fragment thereof or complement of a fragment thereof.

Also provided herein is a method for diagnosing or determining a predisposition for lymphomas, comprising sequencing at least one Nrf2 gene from an individual and determining the nucleic acid sequence of the Nrf2 gene or a fragment thereof. In a preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:210, or a fragment thereof. In another preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:212, or a fragment thereof.

Similarly provided are methods for determining lymphoma subtype and determining a prognosis for an individual having lymphoma, which comprise sequencing at least one Nrf2 gene from an individual and determining the nucleic acid sequence of the Nrf2 gene or a fragment thereof. In a preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:210, or a fragment thereof. In another preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:212, or a fragment thereof.

In yet another aspect of the invention, a method is provided for determining the number of copies of an Nrf2 gene in an individual. In a preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:210, or complement thereof, or a fragment thereof or complement of a fragment thereof. In a preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:212, or complement thereof, or a fragment thereof or complement of a fragment thereof.

In yet another aspect of the invention, a method is provided for determining the chromosomal location of an Nrf2 gene. In a preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:210, or a fragment thereof. In another preferred embodiment, an Nrf2 gene comprises the nucleic acid sequence set forth in SEQ ID NO:212, or a fragment thereof. Such a 5 method may be used to determine Nrf2 gene rearrangements or translocations. Without being bound by theory, Nrf2 gene rearrangement and translocation events appear to be important in the aetiology of lymphoma.

It is an object of this invention that the identification Nrf2 genes and recognition of their involvement in 10 lymphoma provide diagnostic agents to distinguish between lymphoma subtypes, and analytical agents for further analysis of mechanisms involved in dysregulated growth and/or survival and/or apoptosis in cells of the hematopoietic system. An additional object of the invention is to provide appropriate and potentially novel targets for therapeutic interventions, particularly with regard to 15 lymphoma, which are identified through the use of the diagnostic and analytical agents provided herein.

Without being bound by theory, it is recognized herein that the involvement of Nrf2 genes in the 20 cellular dysregulation underlying lymphoma implicates genes having an Nrf2 DNA binding sequence in the cellular dysregulation underlying lymphoma. In a preferred embodiment, the Nrf2 DNA binding sequence is bound by an Nrf2 protein comprising the amino acid sequence set forth in SEQ ID NO:211 and at Genbank accession number AAA68291, or a fragment thereof. In another preferred embodiment, the Nrf2 DNA binding sequence is bound by an Nrf2 protein comprising the amino acid sequence set forth in SEQ ID NO:213 and at Genbank accession number NP_006155, or a fragment thereof.

Novel sequences are also provided herein. Other aspects of the invention will become apparent to the skilled artisan by the following description of the invention.

25 DETAILED DESCRIPTION OF THE INVENTION

The present invention is directed to a number of sequences associated with lymphoma. The use of oncogenic retroviruses, whose sequences insert into the genome of the host organism resulting in 30 lymphoma, allows the identification of host sequences involved in lymphoma. These sequences may then be used in a number of different ways, including diagnosis, prognosis, screening for modulators (including both agonists and antagonists), antibody generation (for immunotherapy and imaging), etc.

Accordingly, the present invention provides nucleic acid and protein sequences that are associated with lymphoma, herein termed "lymphoma/leukemia associated" or "lymphoma/leukemia defining" or "LA" sequences.

In a preferred embodiment, the present invention sets forth LA nucleic acids referred to herein as Pik3r1 nucleic acids. In another preferred embodiment, the present invention sets forth LA proteins referred to herein as Pik3r1 proteins.

5 In addition, the present invention provides GNAS nucleic acid and protein sequences that are associated with lymphoma. Gnas protein sequences include those encoded by a GNAS nucleic acid. Known proteins encoded by GNAS include G_sα, XLα_s and NESP55.

In addition, the present invention provides HIPK1 nucleic acid and protein sequences that are associated with lymphoma.

In a preferred embodiment the LA sequence is JAK1.

10 In a preferred embodiment, the LA sequence is Neurogranin.

In a preferred embodiment, the present invention sets forth LA nucleic acids referred to herein as Nrf2 nucleic acids. In another preferred embodiment, the present invention sets forth LA proteins referred to herein as Nrf2 proteins.

15 "Association" in this context means that the nucleotide or protein sequences are either differentially expressed or altered in lymphoma as compared to normal lymphoid tissue. As outlined below, LA sequences include those that are up-regulated (i.e. expressed at a higher level) in lymphoma, as well as those that are down-regulated (i.e. expressed at a lower level), in lymphoma. LA sequences also include sequences which have been altered (i.e., truncated sequences or sequences with a point mutation) and show either the same expression profile or an altered profile. In a preferred embodiment, the LA sequences are from humans; however, as will be appreciated by those in the art, LA sequences from other organisms may be useful in animal models of disease and drug evaluation; thus, other LA sequences are provided, from vertebrates, including mammals, including rodents (rats, mice, hamsters, guinea pigs, etc.), primates, farm animals (including sheep, goats, pigs, cows, horses, etc). LA sequences from other organisms may be obtained using the techniques outlined 20 below.

25 LA sequences can include both nucleic acid and amino acid sequences. In a preferred embodiment, the LA sequences are recombinant nucleic acids. By the term "recombinant nucleic acid" herein is meant nucleic acid, originally formed in vitro, in general, by the manipulation of nucleic acid by polymerases and endonucleases, in a form not normally found in nature. Thus an isolated nucleic acid, in a linear form, or an expression vector formed in vitro by ligating DNA molecules that are not 30 normally joined, are both considered recombinant for the purposes of this invention. It is understood that once a recombinant nucleic acid is made and reintroduced into a host cell or organism, it will replicate non-recombinantly, i.e. using the in vivo cellular machinery of the host cell rather than in vitro

manipulations; however, such nucleic acids, once produced recombinantly, although subsequently replicated non-recombinantly, are still considered recombinant for the purposes of the invention.

Similarly, a "recombinant protein" is a protein made using recombinant techniques, i.e. through the expression of a recombinant nucleic acid as depicted above. A recombinant protein is distinguished from naturally occurring protein by at least one or more characteristics. For example, the protein may be isolated or purified away from some or all of the proteins and compounds with which it is normally associated in its wild type host, and thus may be substantially pure. For example, an isolated protein is unaccompanied by at least some of the material with which it is normally associated in its natural state, preferably constituting at least about 0.5%, more preferably at least about 5% by weight of the total protein in a given sample. A substantially pure protein comprises at least about 75% by weight of the total protein, with at least about 80% being preferred, and at least about 90% being particularly preferred. The definition includes the production of an LA protein from one organism in a different organism or host cell. Alternatively, the protein may be made at a significantly higher concentration than is normally seen, through the use of an inducible promoter or high expression promoter, such that the protein is made at increased concentration levels. Alternatively, the protein may be in a form not normally found in nature, as in the addition of an epitope tag or amino acid substitutions, insertions and deletions, as discussed below.

In a preferred embodiment, the LA sequences are nucleic acids. As will be appreciated by those in the art and is more fully outlined below, LA sequences are useful in a variety of applications, including diagnostic applications, which will detect naturally occurring nucleic acids, as well as screening applications; for example, biochips comprising nucleic acid probes to the LA sequences can be generated. In the broadest sense, then, by "nucleic acid" or "oligonucleotide" or grammatical equivalents herein means at least two nucleotides covalently linked together. A nucleic acid of the present invention will generally contain phosphodiester bonds, although in some cases, as outlined below (for example in antisense applications or when a candidate agent is a nucleic acid), nucleic acid analogs may be used that have alternate backbones, comprising, for example, phosphoramide (Beaucage et al., Tetrahedron 49(10):1925 (1993) and references therein; Letsinger, J. Org. Chem. 35:3800 (1970); Sprinzel et al., Eur. J. Biochem. 81:579 (1977); Letsinger et al., Nucl. Acids Res. 14:3487 (1986); Sawai et al., Chem. Lett. 805 (1984), Letsinger et al., J. Am. Chem. Soc. 110:4470 (1988); and Pauwels et al., Chemica Scripta 26:141 91986)), phosphorothioate (Mag et al., Nucleic Acids Res. 19:1437 (1991); and U.S. Patent No. 5,644,048), phosphorodithioate (Briu et al., J. Am. Chem. Soc. 111:2321 (1989), O-methylphosphoroamidite linkages (see Eckstein, Oligonucleotides and Analogues: A Practical Approach, Oxford University Press), and peptide nucleic acid backbones and linkages (see Egholm, J. Am. Chem. Soc. 114:1895 (1992); Meier et al., Chem. Int. Ed. Engl. 31:1008 (1992); Nielsen, Nature, 365:566 (1993); Carlsson et al., Nature 380:207 (1996), all of which are incorporated by reference). Other analog nucleic acids include those with positive backbones (Denpcy et al., Proc. Natl. Acad. Sci. USA 92:6097 (1995); non-ionic backbones (U.S. Patent Nos. 5,386,023, 5,637,684, 5,602,240, 5,216,141 and 4,469,863; Kiedrowski et al., Angew. Chem. Intl. Ed. English 30:423 (1991); Letsinger et al., J. Am. Chem. Soc. 110:4470 (1988); Letsinger et al., Nucleoside &

Nucleotide 13:1597 (1994); Chapters 2 and 3, ASC Symposium Series 580, "Carbohydrate Modifications in Antisense Research", Ed. Y.S. Sanghui and P. Dan Cook; Mesmaeker et al., Bioorganic & Medicinal Chem. Lett. 4:395 (1994); Jeffs et al., J. Biomolecular NMR 34:17 (1994); Tetrahedron Lett. 37:743 (1996)) and non-ribose backbones, including those described in U.S. Patent Nos. 5,235,033 and 5,034,506, and Chapters 6 and 7, ASC Symposium Series 580, "Carbohydrate Modifications in Antisense Research", Ed. Y.S. Sanghui and P. Dan Cook. Nucleic acids containing one or more carbocyclic sugars are also included within one definition of nucleic acids (see Jenkins et al., Chem. Soc. Rev. (1995) pp169-176). Several nucleic acid analogs are described in Rawls, C & E News June 2, 1997 page 35. All of these references are hereby expressly incorporated by reference.

5 These modifications of the ribose-phosphate backbone may be done for a variety of reasons, for example to increase the stability and half-life of such molecules in physiological environments or as probes on a biochip.

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As will be appreciated by those in the art, all of these nucleic acid analogs may find use in the present invention. In addition, mixtures of naturally occurring nucleic acids and analogs can be made; 15 alternatively, mixtures of different nucleic acid analogs, and mixtures of naturally occurring nucleic acids and analogs may be made.

Particularly preferred are peptide nucleic acids (PNA) which includes peptide nucleic acid analogs. These backbones are substantially non-ionic under neutral conditions, in contrast to the highly charged phosphodiester backbone of naturally occurring nucleic acids. This results in two 20 advantages. First, the PNA backbone exhibits improved hybridization kinetics. PNAs have larger changes in the melting temperature (T_m) for mismatched versus perfectly matched basepairs. DNA and RNA typically exhibit a 2-4°C drop in T_m for an internal mismatch. With the non-ionic PNA backbone, the drop is closer to 7-9°C. Similarly, due to their non-ionic nature, hybridization of the bases attached to these backbones is relatively insensitive to salt concentration. In addition, PNAs 25 are not degraded by cellular enzymes, and thus can be more stable.

The nucleic acids may be single stranded or double stranded, as specified, or contain portions of both double stranded or single stranded sequence. As will be appreciated by those in the art, the depiction of a single strand ("Watson") also defines the sequence of the other strand ("Crick"); thus the sequences described herein also includes the complement of the sequence. The nucleic acid may be 30 DNA, both genomic and cDNA, RNA or a hybrid, where the nucleic acid contains any combination of deoxyribo- and ribo-nucleotides, and any combination of bases, including uracil, adenine, thymine, cytosine, guanine, inosine, xanthine hypoxanthine, isocytosine, isoguanine, etc. As used herein, the term "nucleoside" includes nucleotides and nucleoside and nucleotide analogs, and modified 35 nucleosides such as amino modified nucleosides. In addition, "nucleoside" includes non-naturally occurring analog structures. Thus for example the individual units of a peptide nucleic acid, each containing a base, are referred to herein as a nucleoside.

An LA sequence can be initially identified by substantial nucleic acid and/or amino acid sequence homology to the LA sequences outlined herein. Such homology can be based upon the overall nucleic acid or amino acid sequence, and is generally determined as outlined below, using either homology programs or hybridization conditions.

5 The LA sequences of the invention were identified as described in the examples; basically, infection of mice with murine leukemia viruses (MuLV; including SL3-3, Akv and mutants thereof) resulted in lymphoma. The LA sequences outlined herein comprise the insertion sites for the virus. In general, the retrovirus can cause lymphoma in three basic ways: first of all, by inserting upstream of a normally silent host gene and activating it (e.g. promoter insertion); secondly, by truncating a host gene that
10 leads to oncogenesis; or by enhancing the transcription of a neighboring gene. By neighboring gene is meant a gene within 100 kb to 500 kb or more, more preferably 50 kb to 100 kb, more preferably 1 kb to 50kb, of the insertion site. For example, retrovirus enhancers, including SL3-3, are known to act on genes up to approximately 200 kilobases of the insertion site.

15 In a preferred embodiment, LA sequences are those that are up-regulated in lymphoma; that is, the expression of these genes is higher in lymphoma as compared to normal lymphoid tissue of the same differentiation stage. "Up-regulation" as used herein means at least about 50%, more preferably at least about 100%, more preferably at least about 150%, more preferably, at least about 200%, with from 300 to at least 1000% being especially preferred.

20 In a preferred embodiment, LA sequences are those that are down-regulated in lymphoma; that is, the expression of these genes is lower in lymphoma as compared to normal lymphoid tissue of the same differentiation stage. "Down-regulation" as used herein means at least about 50%, more preferably at least about 100%, more preferably at least about 150%, more preferably, at least about 200%, with from 300 to at least 1000% being especially preferred.

25 In a preferred embodiment, LA sequences are those that are altered but show either the same expression profile or an altered profile as compared to normal lymphoid tissue of the same differentiation stage. "Altered LA sequences" as used herein refers to sequences which are truncated, contain insertions or contain point mutations.

30 In a preferred embodiment, Pik3r1 sequences are those that are altered but show either the same expression profile or an altered profile as compared to normal lymphoid tissue of the same differentiation stage. "Altered Pik3r1 sequences" as used herein refers to sequences which are truncated, contain insertions, deletions, fusions, or contain point mutations.

35 In one embodiment, the present invention provides an Pik3r1 gene comprising the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank Accession number U50413. In one embodiment, the present invention provides an Pik3r1 gene comprising the nucleic acid sequence set forth by nucleotides 575 to 2749 in SEQ ID NO:178 and at Genbank Accession number U50413.

In one embodiment, the present invention provides an Pik3r1 gene comprising the nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank Accession number M61906. In one embodiment, the present invention provides an Pik3r1 gene comprising the nucleic acid sequence set forth by nucleotides 43 to 2217 in SEQ ID NO:180 and at Genbank Accession number M61906.

5 In one embodiment, the present invention provides a Pik3r1 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank Accession number U50413. In one embodiment, the present invention provides an Pik3r1 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 575 to 2749 in SEQ ID NO:178 and at Genbank Accession number
10 U50413.

In one embodiment, the present invention provides a Pik3r1 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank Accession number M61906. In one embodiment, the present invention provides an Pik3r1 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 43 to 2217 in SEQ ID NO:180 and at Genbank Accession number
15 M61906.

In one embodiment, the present invention provides an Pik3r1 gene comprising a nucleic acid that hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank Accession number U50413.

20 In one embodiment, the present invention provides an Pik3r1 gene comprising a nucleic acid that hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank Accession number M61906.

In one embodiment, the present invention provides an Pik3r1 gene encoding an SH2 domain-containing protein, comprising the nucleic acid sequence set forth by nucleotides 1568-1811, or 1571-
25 1796, or 2444-2666, or 2444-2681 in SEQ ID NO:1 and at Genbank Accession number U50413. In one embodiment, the present invention provides an Pik3r1 gene encoding an SH2 domain-containing protein, comprising a nucleic acid which hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth by nucleotides 1568-1811, or 1571-1796, or 2444-
30 2666, or 2444-2681 in SEQ ID NO:178 and at Genbank Accession number U50413. In one embodiment, the present invention provides an Pik3r1 gene encoding an SH2 domain-containing protein, comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 1568-1811, or 1571-1796, or 2444-2666, or 2444-2681 in SEQ ID NO:178 and at Genbank Accession number U50413.

35 In one embodiment, the present invention provides an Pik3r1 gene encoding an SH3 domain-containing protein, comprising the nucleic acid sequence set forth by nucleotides 4-75, or 7-77 in SEQ

ID NO:178 and at Genbank accession number U50413. In one embodiment, the present invention provides an Pik3r1 gene encoding an SH3 domain-containing protein, comprising a nucleic acid which will hybridize under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth by nucleotides 4-75, or 7-77 in SEQ ID NO:178 and at Genbank accession number U50413.

5 In one embodiment, the present invention provides an Pik3r1 gene encoding an SH3 domain-containing protein, comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 4-75, or 7-77 in SEQ ID NO:178 and at Genbank accession number U50413.

10 In one embodiment, the present invention provides an Pik3r1 gene encoding a protein comprising a RhoGAP domain, comprising the nucleic acid sequence set forth by nucleotides 142-277, or 143-293 in SEQ ID NO:178 and at Genbank accession number U50413. In one embodiment, the present invention provides an Pik3r1 gene encoding a protein comprising a RhoGAP domain, comprising a nucleic acid which will hybridize under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth by nucleotides 142-277, or 143-293 in SEQ ID NO:178 and at Genbank accession number U50413. In one embodiment, the present invention provides an Pik3r1 gene encoding a protein comprising a RhoGAP domain, comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 142-277, or 143-293 in SEQ ID NO:178 and at Genbank accession number U50413.

15 In one embodiment, the present invention provides an Pik3r1 gene encoding an SH2 domain-containing protein, comprising the nucleic acid sequence set forth by nucleotides 1037-1280, or 1913-2150, or 1040-1265, or 1913-3035 in SEQ ID NO:180 and at Genbank Accession number M61906. In one embodiment, the present invention provides an Pik3r1 gene encoding an SH2 domain-containing protein, comprising a nucleic acid which hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth by nucleotides 1037-1280, or 1913-2150, or 1040-25 1265, or 1913-3035 in SEQ ID NO:180 and at Genbank Accession number M61906. In one embodiment, the present invention provides an Pik3r1 gene encoding an SH2 domain-containing protein, comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 1037-1280, or 1913-2150, or 1040-1265, or 1913-3035 in SEQ ID NO:180 and at Genbank Accession number M61906.

20 In one embodiment, the present invention provides an Pik3r1 gene encoding an SH3 domain-containing protein, comprising the nucleic acid sequence set forth by nucleotides 53-266 or 62-272 in SEQ ID NO:180 and at Genbank accession number M61906. In one embodiment, the present invention provides an Pik3r1 gene encoding an SH3 domain-containing protein, comprising a nucleic acid which will hybridize under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth by nucleotides 53-266 or 62-272 in SEQ ID NO:180 and at Genbank accession 35 number M61906. In one embodiment, the present invention provides an Pik3r1 gene encoding an SH3 domain-containing protein, comprising a nucleic acid sequence having at least about 90% identity

to the nucleic acid sequence set forth by nucleotides 53-266 or 62-272 in SEQ ID NO:180 and at Genbank accession number M61906.

5 In one embodiment, the present invention provides an Pik3r1 gene encoding a protein comprising a RhoGAP domain, comprising the nucleic acid sequence set forth by nucleotides 428-929 or 428-872 in SEQ ID NO:180 and at Genbank accession number M61906. In one embodiment, the present invention provides an Pik3r1 gene encoding a protein comprising a RhoGAP domain, comprising a nucleic acid which will hybridize under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth by nucleotides 428-929 or 428-872 in SEQ ID NO:180 and at Genbank accession number M61906. In one embodiment, the present invention provides an Pik3r1 gene 10 encoding a protein comprising a RhoGAP domain, comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 428-929 or 428-872 in SEQ ID NO:180 and at Genbank accession number M61906.

15 In one embodiment, the present invention provides an Pik3r1 gene comprising a nucleic acid sequence that encodes an Pik3r1 protein comprising the amino acid sequence set forth in SEQ ID NO:179 and at Genbank Accession Number AAC52847.

In one embodiment, the present invention provides an Pik3r1 gene comprising a nucleic acid sequence that encodes an Pik3r1 protein comprising the amino acid sequence set forth in SEQ ID NO:181 and at Genbank Accession Number A38748.

20 In one embodiment, the present invention provides an Pik3r1 gene encoding an SH2 domain-containing Pik3r1 protein comprising the amino acid sequence set forth by amino acids 332-413, or 333-408, or 624-703, or 624-698, in SEQ ID NO:179 and at Genbank Accession Number AAC52847.

In one embodiment, the present invention provides an Pik3r1 gene encoding an SH2 domain-containing Pik3r1 protein comprising the amino acid sequence set forth by amino acids 332-413, or 333-408, or 624-703, or 624-698, in SEQ ID NO:181 and at Genbank Accession Number A38748.

25 In one embodiment, the present invention provides an Pik3r1 gene encoding an SH3 domain-containing Pik3r1 protein comprising the amino acid sequence set forth by amino acids 4-75 or 7-77 in SEQ ID NO:179 and at Genbank accession number AAC52847.

30 In one embodiment, the present invention provides an Pik3r1 gene encoding an SH3 domain-containing Pik3r1 protein comprising the amino acid sequence set forth by amino acids 4-75 or 7-77 in SEQ ID NO:181 and at Genbank accession number A38748.

In one embodiment, the present invention provides an Pik3r1 gene encoding RhoGAP domain-containing Pik3r1 protein comprising the amino acid sequence set forth by amino acids 142-277 or 143-293 in SEQ ID NO:179 and at Genbank accession number AAC52847.

In one embodiment, the present invention provides an Pik3r1 gene encoding RhoGAP domain-containing Pik3r1 protein comprising the amino acid sequence set forth by amino acids 129-296 or 129-277 in SEQ ID NO:179 and at Genbank accession number M61906.

- 5 In one embodiment, the present invention provides Pik3r1 proteins encoded by Pik3r1 nucleic acids as described herein.

In a preferred embodiment, the present invention sets forth LA nucleic acids referred to herein as Nrf2 nucleic acids. In another preferred embodiment, the present invention sets forth LA proteins referred to herein as Nrf2 proteins.

- 10 In one embodiment, the present invention provides an Nrf2 gene comprising the nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank Accession number U20532. In one embodiment, the present invention provides an Nrf2 gene comprising the nucleic acid sequence set forth by nucleotides 298 to 2043 in SEQ ID NO:210 and at Genbank Accession number U20532.

- 15 In one embodiment, the present invention provides an Nrf2 gene comprising the nucleic acid sequence set forth in SEQ ID NO:212 and at Genbank Accession number NM_006164. In one embodiment, the present invention provides an Nrf2 gene comprising the nucleic acid sequence set forth by nucleotides 40 to 1809 in SEQ ID NO:212 and at Genbank Accession number NM_006164.

- 20 In one embodiment, the present invention provides a Nrf2 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank Accession number U20532. In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 298 to 2043 in SEQ ID NO:210 and at Genbank Accession number U20532.

- 25 In one embodiment, the present invention provides a Nrf2 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:212 and at Genbank Accession number NM_006164. In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 40 to 1809 in SEQ ID NO:212 and at Genbank Accession number NM_006164.

- 30 In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid that hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank Accession number U20532.

In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid that hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:212 and at Genbank Accession number NM_006164.

5 In one embodiment, the present invention provides an Nrf2 gene comprising the nucleic acid sequence set forth by nucleotides 1716 to 1850 in SEQ ID NO:210 and at Genbank Accession number U20532. In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid which hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth by nucleotides 1716 to 1850 in SEQ ID NO:210 and at Genbank Accession number U20532. In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 1716 to 1850 in SEQ ID NO:210 and at Genbank Accession number U20532.

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15 In one embodiment, the present invention provides an Nrf2 gene comprising the nucleic acid sequence set forth by nucleotides 1482 to 1616, more preferably 1482 to 1550, in SEQ ID NO:212 and at Genbank Accession number NM_006164. In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid which hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth by nucleotides 1482 to 1616, more preferably 1482 to 1550, in SEQ ID NO:212 and at Genbank Accession number NM_006164. In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 1482 to 1616, more preferably 1482 to 1550, in SEQ ID NO:212 and at Genbank Accession number NM_006164.

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In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence that encodes an Nrf2 protein comprising the amino acid sequence set forth in SEQ ID NO:211 and at Genbank Accession Number AAA68291.

25 In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence that encodes an Nrf2 protein comprising the amino acid sequence set forth in SEQ ID NO:213 and at Genbank Accession Number NP_006155.

In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence encoding an Nrf2 protein comprising the amino acid sequence set forth by amino acids 474 to 518 in SEQ ID NO:211 and at Genbank Accession Number AAA68291.

30 In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence encoding an Nrf2 protein comprising the amino acid sequence set forth by amino acids 482 to 526, more preferably 482 to 504, in SEQ ID NO:213 and at Genbank Accession Number NP_006155.

In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence encoding an Nrf2 protein comprising the amino acid sequence set forth in SEQ ID NO:211 and at

Genbank Accession Number AAA68291, except for lacking a fragment of the amino acid sequence set forth by amino acids 474 to 518 in SEQ ID NO:211 and at Genbank Accession Number AAA68291.

5 In one embodiment, the present invention provides an Nrf2 gene comprising a nucleic acid sequence encoding an Nrf2 protein comprising the amino acid sequence set forth in SEQ ID NO:213 and at Genbank Accession Number NP_006155, except for lacking a fragment of the amino acid sequence set forth by amino acids 482 to 526, more preferably 482 to 504, in SEQ ID NO:213 and at Genbank Accession Number NP_006155.

10 In one embodiment, the present invention provides Nrf2 proteins encoded by Nrf2 nucleic acids as described herein.

LA proteins of the present invention may be classified as secreted proteins, transmembrane proteins or intracellular proteins.

15 In a preferred embodiment the LA protein is an intracellular protein. Intracellular proteins may be found in the cytoplasm and/or in the nucleus. Intracellular proteins are involved in all aspects of cellular function and replication (including, for example, signaling pathways); aberrant expression of such proteins results in unregulated or disregulated cellular processes. For example, many intracellular proteins have enzymatic activity such as protein kinase activity, protein phosphatase activity, protease activity, nucleotide cyclase activity, polymerase activity and the like. Intracellular proteins also serve 20 as docking proteins that are involved in organizing complexes of proteins, or targeting proteins to various subcellular localizations, and are involved in maintaining the structural integrity of organelles.

25 In its native form, Pik3r1 protein is an intracellular protein comprising SH2, Sh3, and RhoGAP domains. Intracellular proteins may be found in the cytoplasm and/or in the nucleus. Intracellular proteins are involved in all aspects of cellular function and replication (including, for example, signaling pathways); aberrant expression of such proteins results in unregulated or disregulated cellular processes. For example, many intracellular proteins have enzymatic activity such as protein kinase 30 activity, phosphatidyl inositol-conjugated lipid kinase activity, protein phosphatase activity, phosphatidyl inositol-conjugated lipid phosphatase activity, protease activity, nucleotide cyclase activity, polymerase activity and the like. Intracellular proteins also serve as docking proteins that are involved in organizing complexes of proteins, or targeting proteins to various subcellular localizations, and are involved in maintaining the structural integrity of organelles.

An increasingly appreciated concept in characterizing intracellular proteins is the presence in the proteins of one or more motifs for which defined functions have been attributed. In addition to the highly conserved sequences found in the enzymatic domain of proteins, highly conserved sequences have been identified in proteins that are involved in protein-protein interaction. For example, Src-homology-2 (SH2) domains bind tyrosine-phosphorylated targets in a sequence dependent manner.

PTB domains, which are distinct from SH2 domains, also bind tyrosine phosphorylated targets. SH3 domains bind to proline-rich targets. In addition, PH domains, tetratricopeptide repeats and WD domains to name only a few, have been shown to mediate protein-protein interactions. Some of these may also be involved in binding to phospholipids or other second messengers. As will be appreciated by one of ordinary skill in the art, these motifs can be identified on the basis of primary sequence; thus, an analysis of the sequence of proteins may provide insight into both the enzymatic potential of the molecule and/or molecules with which the protein may associate.

Common protein motifs have also been identified among transcription factors and have been used to divide these factors into families. These motifs include the basic helix-loop-helix, basic leucine zipper, zinc finger and homeodomain motifs.

HIPK1 is known to contain several conserved domains, including a homeoprotein interaction domain, a protein kinase domain, a PEST domain, and a YH domain enriched in tyrosine and histidine residues (Kim et al., J. Biol. Chem. 273:25875 (1998)). In the mouse HIPK1 amino acid sequence depicted in Table 16 as SEQ ID NO. 197, the homeoprotein interaction domain is from about amino acid 190 to about amino acid 518, the protein kinase domain is from about amino acid 581 to about amino acid 848, the PEST domain is from about amino acid 890 to about amino acid 974, and the YH domain is from about amino acid 1067 to about amino acid 1210.

In a preferred embodiment, the LA sequences are transmembrane proteins or can be made to be transmembrane proteins through the use of recombinant DNA technology. Transmembrane proteins are molecules that span the phospholipid bilayer of a cell. They may have an intracellular domain, an extracellular domain, or both. The intracellular domains of such proteins may have a number of functions including those already described for intracellular proteins. For example, the intracellular domain may have enzymatic activity and/or may serve as a binding site for additional proteins. Frequently the intracellular domain of transmembrane proteins serves both roles. For example certain receptor tyrosine kinases have both protein kinase activity and SH2 domains. In addition, autophosphorylation of tyrosines on the receptor molecule itself, creates binding sites for additional SH2 domain containing proteins.

Transmembrane proteins may contain from one to many transmembrane domains. For example, receptor tyrosine kinases, certain cytokine receptors, receptor guanylyl cyclases and receptor serine/threonine protein kinases contain a single transmembrane domain. However, various other proteins including channels and adenylyl cyclases contain numerous transmembrane domains. Many important cell surface receptors are classified as "seven transmembrane domain" proteins, as they contain 7 membrane spanning regions. Important transmembrane protein receptors include, but are not limited to insulin receptor, insulin-like growth factor receptor, human growth hormone receptor, glucose transporters, transferrin receptor, epidermal growth factor receptor, low density lipoprotein receptor, epidermal growth factor receptor, leptin receptor, interleukin receptors, e.g. IL-1 receptor, IL-2 receptor, etc.

Characteristics of transmembrane domains include approximately 20 consecutive hydrophobic amino acids that may be followed by charged amino acids. Therefore, upon analysis of the amino acid sequence of a particular protein, the localization and number of transmembrane domains within the protein may be predicted.

- 5 The extracellular domains of transmembrane proteins are diverse; however, conserved motifs are found repeatedly among various extracellular domains. Conserved structure and/or functions have been ascribed to different extracellular motifs. For example, cytokine receptors are characterized by a cluster of cysteines and a WSXWS (W= tryptophan, S= serine, X=any amino acid) motif.
10 Immunoglobulin-like domains are highly conserved. Mucin-like domains may be involved in cell adhesion and leucine-rich repeats participate in protein-protein interactions.

Many extracellular domains are involved in binding to other molecules. In one aspect, extracellular domains are receptors. Factors that bind the receptor domain include circulating ligands, which may be peptides, proteins, or small molecules such as adenosine and the like. For example, growth factors such as EGF, FGF and PDGF are circulating growth factors that bind to their cognate
15 receptors to initiate a variety of cellular responses. Other factors include cytokines, mitogenic factors, neurotrophic factors and the like. Extracellular domains also bind to cell-associated molecules. In this respect, they mediate cell-cell interactions. Cell-associated ligands can be tethered to the cell for example via a glycosylphosphatidylinositol (GPI) anchor, or may themselves be transmembrane proteins. Extracellular domains also associate with the extracellular matrix and contribute to the
20 maintenance of the cell structure.

LA proteins that are transmembrane are particularly preferred in the present invention as they are good targets for immunotherapeutics, as are described herein. In addition, as outlined below, transmembrane proteins can be also useful in imaging modalities.

It will also be appreciated by those in the art that a transmembrane protein can be made soluble by
25 removing transmembrane sequences, for example through recombinant methods. Furthermore, transmembrane proteins that have been made soluble can be made to be secreted through recombinant means by adding an appropriate signal sequence.

It is further recognized that Nrf2 proteins can be made to be secreted proteins though recombinant methods. Secretion can be either constitutive or regulated. Secreted proteins have a signal peptide
30 or signal sequence that targets the molecule to the secretory pathway.

In another preferred embodiment, the Nrf2 proteins are nuclear proteins, preferably transcription factors. Transcription factors are involved in numerous physiological events and act by regulating gene expression at the transcriptional level. Transcription factors often serve as nodal points of regulation controlling multiple genes. They are capable of effecting a multifarious change in gene
35 expression and can integrate many convergent signals to effect such a change. Transcription factors

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are often regarded as "master regulators" of a particular cellular state or event. Accordingly, transcription factors have often been found to faithfully mark a particular cell state, a quality which makes them attractive for use as diagnostic markers. In addition, because of their important role as coordinators of patterns of gene expression associated with particular cell states, transcription factors are attractive therapeutic targets. Intervention at the level of transcriptional regulation allows one to effectively target multiple genes associated with a dysfunction which fall under the regulation of a "master regulator" or transcription factor.

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In a preferred embodiment, the LA proteins are secreted proteins; the secretion of which can be either constitutive or regulated. These proteins have a signal peptide or signal sequence that targets the molecule to the secretory pathway. Secreted proteins are involved in numerous physiological events; by virtue of their circulating nature, they serve to transmit signals to various other cell types. The secreted protein may function in an autocrine manner (acting on the cell that secreted the factor), a paracrine manner (acting on cells in close proximity to the cell that secreted the factor) or an endocrine manner (acting on cells at a distance). Thus secreted molecules find use in modulating or altering numerous aspects of physiology. LA proteins that are secreted proteins are particularly preferred in the present invention as they serve as good targets for diagnostic markers, for example for blood tests.

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An LA sequence is initially identified by substantial nucleic acid and/or amino acid sequence homology to the LA sequences outlined herein. Such homology can be based upon the overall nucleic acid or amino acid sequence, and is generally determined as outlined below, using either homology programs or hybridization conditions.

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In one embodiment, an Pik3r1 sequence can be identified by substantial nucleic acid sequence identity or homology to the Pik3r1 nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank Accession number U50413.

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In another embodiment, an Pik3r1 sequence can be identified by substantial nucleic acid sequence identity or homolgy to the Pik3r1 nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank Accession number M61906.

In one embodiment, an Pik3r1 sequence can be identified by substantial amino acid sequence identity or homology to the Pik3r1 amino acid sequence set forth in SEQ ID NO:179 and at Genbank Accession number AAC52847.

In another embodiment, an Pik3r1 sequence can be identified by substantial amino acid sequence identity or homology to the Pik3r1 amino acid sequence set forth in SEQ ID NO:181 and at Genbank Accession number A38478.

In one embodiment, an Nrf2 sequence can be identified by substantial nucleic acid sequence identity or homology to the Nrf2 nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank Accession number U20532.

5 In another embodiment, an Nrf2 sequence can be identified by substantial nucleic acid sequence identity or homology to the Nrf2 nucleic acid sequence set forth in SEQ ID NO:210 and at Genbank Accession number NM_006164.

In one embodiment, an Nrf2 sequence can be identified by substantial amino acid sequence identity or homology to the Nrf2 amino acid sequence set forth in SEQ ID NO:211 and at Genbank Accession number AAA68291.

10 In another embodiment, an Nrf2 sequence can be identified by substantial amino acid sequence identity or homology to the Nrf2 amino acid sequence set forth in SEQ ID NO:213 and at Genbank Accession number NP_006155.

As used herein, a nucleic acid is a "LA nucleic acid" if the overall homology of the nucleic acid sequence to one of the nucleic acids of Tables 1, 2, 4, 6, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 15 22, 23, 24, 27, 28 or 30 is preferably greater than about 75%, more preferably greater than about 80%, even more preferably greater than about 85% and most preferably greater than 90%. In some embodiments the homology will be as high as about 93 to 95 or 98%. In a preferred embodiment, the sequences which are used to determine sequence identity or similarity are selected from those of the nucleic acids of Tables 1, 2, 4, 6, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 22, 23, 24, 27, 28 or 30. 20 In another embodiment, the sequences are naturally occurring allelic variants of the sequences of the nucleic acids of Table 1, 2, 3, 4, 6, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 22, 23, 24, 27, 28 or 30 . In another embodiment, the sequences are sequence variants as further described herein.

Homology in this context means sequence similarity or identity, with identity being preferred. A preferred comparison for homology purposes is to compare the sequence containing sequencing errors to the correct sequence. This homology will be determined using standard techniques known in the art, including, but not limited to, the local homology algorithm of Smith & Waterman, Adv. Appl. Math. 2:482 (1981), by the homology alignment algorithm of Needleman & Wunsch, J. Mol. Biol. 48:443 (1970), by the search for similarity method of Pearson & Lipman, PNAS USA 85:2444 (1988), by computerized implementations of these algorithms (GAP, BESTFIT, FASTA, and TFASTA in the 30 Wisconsin Genetics Software Package, Genetics Computer Group, 575 Science Drive, Madison, WI), the Best Fit sequence program described by Devereux et al., Nucl. Acid Res. 12:387-395 (1984), preferably using the default settings, or by inspection.

One example of a useful algorithm is PILEUP. PILEUP creates a multiple sequence alignment from a group of related sequences using progressive, pairwise alignments. It can also plot a tree showing the clustering relationships used to create the alignment. PILEUP uses a simplification of the progressive 35

alignment method of Feng & Doolittle, J. Mol. Evol. 35:351-360 (1987); the method is similar to that described by Higgins & Sharp CABIOS 5:151-153 (1989). Useful PILEUP parameters including a default gap weight of 3.00, a default gap length weight of 0.10, and weighted end gaps.

Another example of a useful algorithm is the BLAST algorithm, described in Altschul et al., J. Mol. Biol. 215, 403-410, (1990) and Karlin et al., PNAS USA 90:5873-5787 (1993). A particularly useful BLAST program is the WU-BLAST-2 program which was obtained from Altschul et al., Methods in Enzymology, 266: 460-480 (1996); <http://blast.wustl>. WU-BLAST-2 uses several search parameters, most of which are set to the default values. The adjustable parameters are set with the following values: overlap span =1, overlap fraction = 0.125, word threshold (T) = 11. The HSP S and HSP S2 parameters are dynamic values and are established by the program itself depending upon the composition of the particular sequence and composition of the particular database against which the sequence of interest is being searched; however, the values may be adjusted to increase sensitivity. A % amino acid sequence identity value is determined by the number of matching identical residues divided by the total number of residues of the "longer" sequence in the aligned region. The "longer" sequence is the one having the most actual residues in the aligned region (gaps introduced by WU-Blast-2 to maximize the alignment score are ignored).

Thus, "percent (%) nucleic acid sequence identity" is defined as the percentage of nucleotide residues in a candidate sequence that are identical with the nucleotide residues of the nucleic acids of the SEQ ID NOS. A preferred method utilizes the BLASTN module of WU-BLAST-2 set to the default parameters, with overlap span and overlap fraction set to 1 and 0.125, respectively.

The alignment may include the introduction of gaps in the sequences to be aligned. In addition, for sequences which contain either more or fewer nucleotides than those of the nucleic acids of the SEQ ID NOS, it is understood that the percentage of homology will be determined based on the number of homologous nucleosides in relation to the total number of nucleosides. Thus, for example, homology of sequences shorter than those of the sequences identified herein and as discussed below, will be determined using the number of nucleosides in the shorter sequence.

In one embodiment, the nucleic acid homology is determined through hybridization studies. Thus, for example, nucleic acids which hybridize under high stringency to the nucleic acids identified in the figures, or their complements, are considered LA sequences. High stringency conditions are known in the art; see for example Maniatis et al., Molecular Cloning: A Laboratory Manual, 2d Edition, 1989, and Short Protocols in Molecular Biology, ed. Ausubel, et al., both of which are hereby incorporated by reference. Stringent conditions are sequence-dependent and will be different in different circumstances. Longer sequences hybridize specifically at higher temperatures. An extensive guide to the hybridization of nucleic acids is found in Tijssen, Techniques in Biochemistry and Molecular Biology--Hybridization with Nucleic Acid Probes, "Overview of principles of hybridization and the strategy of nucleic acid assays" (1993). Generally, stringent conditions are selected to be about 5-10°C lower than the thermal melting point (Tm) for the specific sequence at a defined ionic strength

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pH. The Tm is the temperature (under defined ionic strength, pH and nucleic acid concentration) at which 50% of the probes complementary to the target hybridize to the target sequence at equilibrium (as the target sequences are present in excess, at Tm, 50% of the probes are occupied at equilibrium). Stringent conditions will be those in which the salt concentration is less than about 1.0 M sodium ion, typically about 0.01 to 1.0 M sodium ion concentration (or other salts) at pH 7.0 to 8.3 and the temperature is at least about 30°C for short probes (e.g. 10 to 50 nucleotides) and at least about 60°C for long probes (e.g. greater than 50 nucleotides). Stringent conditions may also be achieved with the addition of destabilizing agents such as formamide.

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In another embodiment, less stringent hybridization conditions are used; for example, moderate or low stringency conditions may be used, as are known in the art; see Maniatis and Ausubel, *supra*, and Tijssen, *supra*.

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In addition, the LA nucleic acid sequences of the invention are fragments of larger genes, i.e. they are nucleic acid segments. Alternatively, the LA nucleic acid sequences can serve as indicators of oncogene position, for example, the LA sequence may be an enhancer that activates a protooncogene. "Genes" in this context includes coding regions, non-coding regions, and mixtures of coding and non-coding regions. Accordingly, as will be appreciated by those in the art, using the sequences provided herein, additional sequences of the LA genes can be obtained, using techniques well known in the art for cloning either longer sequences or the full length sequences; see Maniatis et al., and Ausubel, et al., *supra*, hereby expressly incorporated by reference. In general, this is done using PCR, for example, kinetic PCR.

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Once the LA nucleic acid is identified, it can be cloned and, if necessary, its constituent parts recombined to form the entire LA nucleic acid. Once isolated from its natural source, e.g., contained within a plasmid or other vector or excised therefrom as a linear nucleic acid segment, the recombinant LA nucleic acid can be further used as a probe to identify and isolate other LA nucleic acids, for example additional coding regions. It can also be used as a "precursor" nucleic acid to make modified or variant LA nucleic acids and proteins.

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The LA nucleic acids of the present invention are used in several ways. In a first embodiment, nucleic acid probes to the LA nucleic acids are made and attached to biochips to be used in screening and diagnostic methods, as outlined below, or for administration, for example for gene therapy and/or antisense applications. Alternatively, the LA nucleic acids that include coding regions of LA proteins can be put into expression vectors for the expression of LA proteins, again either for screening purposes or for administration to a patient.

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In a preferred embodiment, nucleic acid probes to LA nucleic acids (both the nucleic acid sequences outlined in the figures and/or the complements thereof) are made. The nucleic acid probes attached to the biochip are designed to be substantially complementary to the LA nucleic acids, i.e. the target sequence (either the target sequence of the sample or to other probe sequences, for example in

sandwich assays), such that hybridization of the target sequence and the probes of the present invention occurs. As outlined below, this complementarity need not be perfect; there may be any number of base pair mismatches which will interfere with hybridization between the target sequence and the single stranded nucleic acids of the present invention. However, if the number of mutations is so great that no hybridization can occur under even the least stringent of hybridization conditions, the sequence is not a complementary target sequence. Thus, by "substantially complementary" herein is meant that the probes are sufficiently complementary to the target sequences to hybridize under normal reaction conditions, particularly high stringency conditions, as outlined herein.

A nucleic acid probe is generally single stranded but can be partially single and partially double stranded. The strandedness of the probe is dictated by the structure, composition, and properties of the target sequence. In general, the nucleic acid probes range from about 8 to about 100 bases long, with from about 10 to about 80 bases being preferred, and from about 30 to about 50 bases being particularly preferred. That is, generally whole genes are not used. In some embodiments, much longer nucleic acids can be used, up to hundreds of bases.

In a preferred embodiment, more than one probe per sequence is used, with either overlapping probes or probes to different sections of the target being used. That is, two, three, four or more probes, with three being preferred, are used to build in a redundancy for a particular target. The probes can be overlapping (i.e. have some sequence in common), or separate.

As will be appreciated by those in the art, nucleic acids can be attached or immobilized to a solid support in a wide variety of ways. By "immobilized" and grammatical equivalents herein is meant the association or binding between the nucleic acid probe and the solid support is sufficient to be stable under the conditions of binding, washing, analysis, and removal as outlined below. The binding can be covalent or non-covalent. By "non-covalent binding" and grammatical equivalents herein is meant one or more of either electrostatic, hydrophilic, and hydrophobic interactions. Included in non-covalent binding is the covalent attachment of a molecule, such as, streptavidin to the support and the non-covalent binding of the biotinylated probe to the streptavidin. By "covalent binding" and grammatical equivalents herein is meant that the two moieties, the solid support and the probe, are attached by at least one bond, including sigma bonds, pi bonds and coordination bonds. Covalent bonds can be formed directly between the probe and the solid support or can be formed by a cross linker or by inclusion of a specific reactive group on either the solid support or the probe or both molecules. Immobilization may also involve a combination of covalent and non-covalent interactions.

In general, the probes are attached to the biochip in a wide variety of ways, as will be appreciated by those in the art. As described herein, the nucleic acids can either be synthesized first, with subsequent attachment to the biochip, or can be directly synthesized on the biochip.

The biochip comprises a suitable solid substrate. By "substrate" or "solid support" or other grammatical equivalents herein is meant any material that can be modified to contain discrete

individual sites appropriate for the attachment or association of the nucleic acid probes and is amenable to at least one detection method. As will be appreciated by those in the art, the number of possible substrates are very large, and include, but are not limited to, glass and modified or functionalized glass, plastics (including acrylics, polystyrene and copolymers of styrene and other materials, polypropylene, polyethylene, polybutylene, polyurethanes, TeflonJ, etc.), polysaccharides, nylon or nitrocellulose, resins, silica or silica-based materials including silicon and modified silicon, carbon, metals, inorganic glasses, etc. In general, the substrates allow optical detection and do not appreciably fluoresce.

In a preferred embodiment, the surface of the biochip and the probe may be derivatized with chemical functional groups for subsequent attachment of the two. Thus, for example, the biochip is derivatized with a chemical functional group including, but not limited to, amino groups, carboxy groups, oxo groups and thiol groups, with amino groups being particularly preferred. Using these functional groups, the probes can be attached using functional groups on the probes. For example, nucleic acids containing amino groups can be attached to surfaces comprising amino groups, for example using linkers as are known in the art; for example, homo- or hetero-bifunctional linkers as are well known (see 1994 Pierce Chemical Company catalog, technical section on cross-linkers, pages 155-200, incorporated herein by reference). In addition, in some cases, additional linkers, such as alkyl groups (including substituted and heteroalkyl groups) may be used.

In this embodiment, the oligonucleotides are synthesized as is known in the art, and then attached to the surface of the solid support. As will be appreciated by those skilled in the art, either the 5' or 3' terminus may be attached to the solid support, or attachment may be via an internal nucleoside.

In an additional embodiment, the immobilization to the solid support may be very strong, yet non-covalent. For example, biotinylated oligonucleotides can be made, which bind to surfaces covalently coated with streptavidin, resulting in attachment.

Alternatively, the oligonucleotides may be synthesized on the surface, as is known in the art. For example, photoactivation techniques utilizing photopolymerization compounds and techniques are used. In a preferred embodiment, the nucleic acids can be synthesized *in situ*, using well known photolithographic techniques, such as those described in WO 95/25116; WO 95/35505; U.S. Patent Nos. 5,700,637 and 5,445,934; and references cited within, all of which are expressly incorporated by reference; these methods of attachment form the basis of the Affimetrix GeneChip™ technology.

In addition to the solid-phase technology represented by biochip arrays, gene expression can also be quantified using liquid-phase arrays. One such system is kinetic polymerase chain reaction (PCR). Kinetic PCR allows for the simultaneous amplification and quantification of specific nucleic acid sequences. The specificity is derived from synthetic oligonucleotide primers designed to preferentially adhere to single-stranded nucleic acid sequences bracketing the target site. This pair of

oligonucleotide primers form specific, non-covalently bound complexes on each strand of the target sequence. These complexes facilitate *in vitro* transcription of double-stranded DNA in opposite orientations. Temperature cycling of the reaction mixture creates a continuous cycle of primer binding, transcription, and re-melting of the nucleic acid to individual strands. The result is an exponential increase of the target dsDNA product. This product can be quantified in real time either through the use of an intercalating dye or a sequence specific probe. SYBR® Greene I, is an example of an intercalating dye, that preferentially binds to dsDNA resulting in a concomitant increase in the fluorescent signal. Sequence specific probes, such as used with TaqMan® technology, consist of a fluorochrome and a quenching molecule covalently bound to opposite ends of an oligonucleotide. The probe is designed to selectively bind the target DNA sequence between the two primers. When the DNA strands are synthesized during the PCR reaction, the fluorochrome is cleaved from the probe by the exonuclease activity of the polymerase resulting in signal dequenching. The probe signaling method can be more specific than the intercalating dye method, but in each case, signal strength is proportional to the dsDNA product produced. Each type of quantification method can be used in multi-well liquid phase arrays with each well representing primers and/or probes specific to nucleic acid sequences of interest. When used with messenger RNA preparations of tissues or cell lines, and an array of probe/primer reactions can simultaneously quantify the expression of multiple gene products of interest. See Germer, S., et al., Genome Res. 10:258-266 (2000); Heid, C. A., et al., Genome Res. 6, 986-994 (1996).

In a preferred embodiment, LA nucleic acids encoding LA proteins are used to make a variety of expression vectors to express LA proteins which can then be used in screening assays, as described below. The expression vectors may be either self-replicating extrachromosomal vectors or vectors which integrate into a host genome. Generally, these expression vectors include transcriptional and translational regulatory nucleic acid operably linked to the nucleic acid encoding the LA protein. The term "control sequences" refers to DNA sequences necessary for the expression of an operably linked coding sequence in a particular host organism. The control sequences that are suitable for prokaryotes, for example, include a promoter, optionally an operator sequence, and a ribosome binding site. Eukaryotic cells are known to utilize promoters, polyadenylation signals, and enhancers.

Nucleic acid is "operably linked" when it is placed into a functional relationship with another nucleic acid sequence. For example, DNA for a presequence or secretory leader is operably linked to DNA for a polypeptide if it is expressed as a preprotein that participates in the secretion of the polypeptide; a promoter or enhancer is operably linked to a coding sequence if it affects the transcription of the sequence; or a ribosome binding site is operably linked to a coding sequence if it is positioned so as to facilitate translation. Generally, "operably linked" means that the DNA sequences being linked are contiguous, and, in the case of a secretory leader, contiguous and in reading phase. However, enhancers do not have to be contiguous. Linking is accomplished by ligation at convenient restriction sites. If such sites do not exist, synthetic oligonucleotide adaptors or linkers are used in accordance with conventional practice. The transcriptional and translational regulatory nucleic acid will generally be appropriate to the host cell used to express the LA protein; for example, transcriptional and

translational regulatory nucleic acid sequences from *Bacillus* are preferably used to express the LA protein in *Bacillus*. Numerous types of appropriate expression vectors, and suitable regulatory sequences are known in the art for a variety of host cells.

5 In general, the transcriptional and translational regulatory sequences may include, but are not limited to, promoter sequences, ribosomal binding sites, transcriptional start and stop sequences, translational start and stop sequences, and enhancer or activator sequences. In a preferred embodiment, the regulatory sequences include a promoter and transcriptional start and stop sequences.

10 Promoter sequences encode either constitutive or inducible promoters. The promoters may be either naturally occurring promoters or hybrid promoters. Hybrid promoters, which combine elements of more than one promoter, are also known in the art, and are useful in the present invention.

15 In addition, the expression vector may comprise additional elements. For example, the expression vector may have two replication systems, thus allowing it to be maintained in two organisms, for example in mammalian or insect cells for expression and in a prokaryotic host for cloning and amplification. Furthermore, for integrating expression vectors, the expression vector contains at least one sequence homologous to the host cell genome, and preferably two homologous sequences which flank the expression construct. The integrating vector may be directed to a specific locus in the host cell by selecting the appropriate homologous sequence for inclusion in the vector. Constructs for integrating vectors are well known in the art.

20 In addition, in a preferred embodiment, the expression vector contains a selectable marker gene to allow the selection of transformed host cells. Selection genes are well known in the art and will vary with the host cell used.

25 The LA proteins of the present invention are produced by culturing a host cell transformed with an expression vector containing nucleic acid encoding an LA protein, under the appropriate conditions to induce or cause expression of the LA protein. The conditions appropriate for LA protein expression will vary with the choice of the expression vector and the host cell, and will be easily ascertained by one skilled in the art through routine experimentation. For example, the use of constitutive promoters in the expression vector will require optimizing the growth and proliferation of the host cell, while the use of an inducible promoter requires the appropriate growth conditions for induction. In addition, in some embodiments, the timing of the harvest is important. For example, the baculoviral systems used in insect cell expression are lytic viruses, and thus harvest time selection can be crucial for product yield.

Appropriate host cells include yeast, bacteria, archaeabacteria, fungi, and insect, plant and animal cells, including mammalian cells. Of particular interest are *Drosophila melanogaster* cells, *Saccharomyces*

cerevisiae and other yeasts, *E. coli*, *Bacillus subtilis*, Sf9 cells, C129 cells, 293 cells, *Neurospora*, BHK, CHO, COS, HeLa cells, THP1 cell line (a macrophage cell line) and human cells and cell lines.

In a preferred embodiment, the LA proteins are expressed in mammalian cells. Mammalian expression systems are also known in the art, and include retroviral systems. A preferred expression vector system is a retroviral vector system such as is generally described in PCT/US97/01019 and PCT/US97/01048, both of which are hereby expressly incorporated by reference. Of particular use as mammalian promoters are the promoters from mammalian viral genes, since the viral genes are often highly expressed and have a broad host range. Examples include the SV40 early promoter, mouse mammary tumor virus LTR promoter, adenovirus major late promoter, herpes simplex virus promoter, and the CMV promoter. Typically, transcription termination and polyadenylation sequences recognized by mammalian cells are regulatory regions located 3' to the translation stop codon and thus, together with the promoter elements, flank the coding sequence. Examples of transcription terminator and polyadenylation signals include those derived from SV40. The methods of introducing exogenous nucleic acid into mammalian hosts, as well as other hosts, is well known in the art, and will vary with the host cell used. Techniques include dextran-mediated transfection, calcium phosphate precipitation, polybrene mediated transfection, protoplast fusion, electroporation, viral infection, encapsulation of the polynucleotide(s) in liposomes, and direct microinjection of the DNA into nuclei.

In a preferred embodiment, LA proteins are expressed in bacterial systems. Bacterial expression systems are well known in the art. Promoters from bacteriophage may also be used and are known in the art. In addition, synthetic promoters and hybrid promoters are also useful; for example, the tac promoter is a hybrid of the trp and lac promoter sequences. Furthermore, a bacterial promoter can include naturally occurring promoters of non-bacterial origin that have the ability to bind bacterial RNA polymerase and initiate transcription. In addition to a functioning promoter sequence, an efficient ribosome binding site is desirable. The expression vector may also include a signal peptide sequence that provides for secretion of the LA protein in bacteria. The protein is either secreted into the growth media (gram-positive bacteria) or into the periplasmic space, located between the inner and outer membrane of the cell (gram-negative bacteria). The bacterial expression vector may also include a selectable marker gene to allow for the selection of bacterial strains that have been transformed. Suitable selection genes include genes which render the bacteria resistant to drugs such as ampicillin, chloramphenicol, erythromycin, kanamycin, neomycin and tetracycline. Selectable markers also include biosynthetic genes, such as those in the histidine, tryptophan and leucine biosynthetic pathways. These components are assembled into expression vectors. Expression vectors for bacteria are well known in the art, and include vectors for *Bacillus subtilis*, *E. coli*, *Streptococcus cremoris*, and *Streptococcus lividans*, among others. The bacterial expression vectors are transformed into bacterial host cells using techniques well known in the art, such as calcium chloride treatment, electroporation, and others.

In one embodiment, LA proteins are produced in insect cells. Expression vectors for the transformation of insect cells, and in particular, baculovirus-based expression vectors, are well known in the art.

5 In a preferred embodiment, LA protein is produced in yeast cells. Yeast expression systems are well known in the art, and include expression vectors for *Saccharomyces cerevisiae*, *Candida albicans* and *C. maltosa*, *Hansenula polymorpha*, *Kluyveromyces fragilis* and *K. lactis*, *Pichia guillermondi* and *P. pastoris*, *Schizosaccharomyces pombe*, and *Yarrowia lipolytica*.

10 The LA protein may also be made as a fusion protein, using techniques well known in the art. Thus, for example, for the creation of monoclonal antibodies. If the desired epitope is small, the LA protein may be fused to a carrier protein to form an immunogen. Alternatively, the LA protein may be made as a fusion protein to increase expression, or for other reasons. For example, when the LA protein is an LA peptide, the nucleic acid encoding the peptide may be linked to other nucleic acid for expression purposes.

15 In one embodiment, the LA nucleic acids, proteins and antibodies of the invention are labeled. By "labeled" herein is meant that a compound has at least one element, isotope or chemical compound attached to enable the detection of the compound. In general, labels fall into three classes: a) isotopic labels, which may be radioactive or heavy isotopes; b) immune labels, which may be antibodies or antigens; and c) colored or fluorescent dyes. The labels may be incorporated into the LA nucleic acids, proteins and antibodies at any position. For example, the label should be capable of producing, 20 either directly or indirectly, a detectable signal. The detectable moiety may be a radioisotope, such as ^3H , ^{14}C , ^{32}P , ^{35}S , or ^{125}I , a fluorescent or chemiluminescent compound, such as fluorescein isothiocyanate, rhodamine, or luciferin, or an enzyme, such as alkaline phosphatase, beta-galactosidase or horseradish peroxidase. Any method known in the art for conjugating the antibody to the label may be employed, including those methods described by Hunter et al., *Nature*, 144:945 25 (1962); David et al., *Biochemistry*, 13:1014 (1974); Pain et al., *J. Immunol. Meth.*, 40:219 (1981); and Nygren, *J. Histochem. and Cytochem.*, 30:407 (1982).

30 Accordingly, the present invention also provides LA protein sequences. An LA protein of the present invention may be identified in several ways. "Protein" in this sense includes proteins, polypeptides, and peptides. As will be appreciated by those in the art, the nucleic acid sequences of the invention can be used to generate protein sequences. There are a variety of ways to do this, including cloning the entire gene and verifying its frame and amino acid sequence, or by comparing it to known 35 sequences to search for homology to provide a frame, assuming the LA protein has homology to some protein in the database being used. Generally, the nucleic acid sequences are input into a program that will search all three frames for homology. This is done in a preferred embodiment using the following NCBI Advanced BLAST parameters. The program is blastx or blastn. The database is nr. The input data is as "Sequence in FASTA format". The organism list is "none". The "expect" is

10; the filter is default. The "descriptions" is 500, the "alignments" is 500, and the "alignment view" is pairwise. The "Query Genetic Codes" is standard (1). The matrix is BLOSUM62; gap existence cost is 11, per residue gap cost is 1; and the lambda ratio is .85 default. This results in the generation of a putative protein sequence.

5 Also included within one embodiment of LA proteins are amino acid variants of the naturally occurring sequences, as determined herein. Preferably, the variants are preferably greater than about 75% homologous to the wild-type sequence, more preferably greater than about 80%, even more preferably greater than about 85% and most preferably greater than 90%. In some embodiments the homology will be as high as about 93 to 95 or 98%. As for nucleic acids, homology in this context means sequence similarity or identity, with identity being preferred. This homology will be determined using 10 standard techniques known in the art as are outlined above for the nucleic acid homologies.

LA proteins of the present invention may be shorter or longer than the wild type amino acid sequences. Thus, in a preferred embodiment, included within the definition of LA proteins are portions or fragments of the wild type sequences herein. In addition, as outlined above, the LA nucleic acids of 15 the invention may be used to obtain additional coding regions, and thus additional protein sequence, using techniques known in the art.

In a preferred embodiment, the LA proteins are derivative or variant LA proteins as compared to the wild-type sequence. That is, as outlined more fully below, the derivative LA peptide will contain at least one amino acid substitution, deletion or insertion, with amino acid substitutions being particularly 20 preferred. The amino acid substitution, insertion or deletion may occur at any residue within the LA peptide.

Also included in an embodiment of LA proteins of the present invention are amino acid sequence variants. These variants fall into one or more of three classes: substitutional, insertional or deletional 25 variants. These variants ordinarily are prepared by site specific mutagenesis of nucleotides in the DNA encoding the LA protein, using cassette or PCR mutagenesis or other techniques well known in the art, to produce DNA encoding the variant, and thereafter expressing the DNA in recombinant cell culture as outlined above. However, variant LA protein fragments having up to about 100-150 residues may be prepared by *in vitro* synthesis using established techniques. Amino acid sequence 30 variants are characterized by the predetermined nature of the variation, a feature that sets them apart from naturally occurring allelic or interspecies variation of the LA protein amino acid sequence. The variants typically exhibit the same qualitative biological activity as the naturally occurring analogue, although variants can also be selected which have modified characteristics as will be more fully outlined below.

While the site or region for introducing an amino acid sequence variation is predetermined, the 35 mutation per se need not be predetermined. For example, in order to optimize the performance of a mutation at a given site, random mutagenesis may be conducted at the target codon or region and the

expressed LA variants screened for the optimal combination of desired activity. Techniques for making substitution mutations at predetermined sites in DNA having a known sequence are well known, for example, M13 primer mutagenesis and LAR mutagenesis. Screening of the mutants is done using assays of LA protein activities.

- 5 Amino acid substitutions are typically of single residues; insertions usually will be on the order of from about 1 to 20 amino acids, although considerably larger insertions may be tolerated. Deletions range from about 1 to about 20 residues, although in some cases deletions may be much larger.

Substitutions, deletions, insertions or any combination thereof may be used to arrive at a final derivative. Generally these changes are done on a few amino acids to minimize the alteration of the 10 molecule. However, larger changes may be tolerated in certain circumstances. When small alterations in the characteristics of the LA protein are desired, substitutions are generally made in accordance with the following chart:

Chart I
Exemplary Substitutions

	Original Residue	Exemplary Substitutions
15	Ala	Ser
	Arg	Lys
	Asn	Gln, His
	Asp	Glu
20	Cys	Ser
	Gln	Asn
	Glu	Asp
	Gly	Pro
	His	Asn, Gln
	Ile	Leu, Val
25	Leu	Ile, Val
	Lys	Arg, Gln, Glu
	Met	Leu, Ile
	Phe	Met, Leu, Tyr
	Ser	Thr
30	Thr	Ser
	Trp	Tyr
	Tyr	Trp, Phe
	Val	Ile, Leu

Substantial changes in function or immunological identity are made by selecting substitutions that are less conservative than those shown in Chart I. For example, substitutions may be made which more significantly affect: the structure of the polypeptide backbone in the area of the alteration, for example the alpha-helical or beta-sheet structure; the charge or hydrophobicity of the molecule at the target site; or the bulk of the side chain. The substitutions which in general are expected to produce the greatest changes in the polypeptide's properties are those in which (a) a hydrophilic residue, e.g. seryl or threonyl is substituted for (or by) a hydrophobic residue, e.g. leucyl, isoleucyl, phenylalanyl, valyl or alanyl; (b) a cysteine or proline is substituted for (or by) any other residue; (c) a residue having an electropositive side chain, e.g. lysyl, arginyl, or histidyl, is substituted for (or by) an electronegative 40 residue.

residue, e.g. glutamyl or aspartyl; or (d) a residue having a bulky side chain, e.g. phenylalanine, is substituted for (or by) one not having a side chain, e.g. glycine.

The variants typically exhibit the same qualitative biological activity and will elicit the same immune response as the naturally-occurring analogue, although variants also are selected to modify the characteristics of the LA proteins as needed. Alternatively, the variant may be designed such that the biological activity of the LA protein is altered. For example, glycosylation sites may be altered or removed, dominant negative mutations created, etc.

Covalent modifications of LA polypeptides are included within the scope of this invention, for example for use in screening. One type of covalent modification includes reacting targeted amino acid residues of an LA polypeptide with an organic derivatizing agent that is capable of reacting with selected side chains or the N- or C-terminal residues of an LA polypeptide. Derivatization with bifunctional agents is useful, for instance, for crosslinking LA to a water-insoluble support matrix or surface for use in the method for purifying anti-LA antibodies or screening assays, as is more fully described below. Commonly used crosslinking agents include, e.g., 1,1-bis(diazoacetyl)-2-phenylethane, glutaraldehyde, N-hydroxysuccinimide esters, for example, esters with 4-azidosalicylic acid, 15 homobifunctional imidoesters, including disuccinimidyl esters such as 3,3'-dithiobis(succinimidylpropionate), bifunctional maleimides such as bis-N-maleimido-1,8-octane and agents such as methyl-3-[(p-azidophenyl)dithio]propioimidate.

Other modifications include deamidation of glutaminyl and asparaginyl residues to the corresponding glutamyl and aspartyl residues, respectively, hydroxylation of proline and lysine, phosphorylation of hydroxyl groups of seryl, threonyl or tyrosyl residues, methylation of the α -amino groups of lysine, arginine, and histidine side chains [T.E. Creighton, Proteins: Structure and Molecular Properties, W.H. Freeman & Co., San Francisco, pp. 79-86 (1983)], acetylation of the N-terminal amine, and amidation of any C-terminal carboxyl group.

Another type of covalent modification of the LA polypeptide included within the scope of this invention comprises altering the native glycosylation pattern of the polypeptide. "Altering the native glycosylation pattern" is intended for purposes herein to mean deleting one or more carbohydrate moieties found in native sequence LA polypeptide, and/or adding one or more glycosylation sites that are not present in the native sequence LA polypeptide.

Addition of glycosylation sites to LA polypeptides may be accomplished by altering the amino acid sequence thereof. The alteration may be made, for example, by the addition of, or substitution by, one or more serine or threonine residues to the native sequence LA polypeptide (for O-linked glycosylation sites). The LA amino acid sequence may optionally be altered through changes at the DNA level, particularly by mutating the DNA encoding the LA polypeptide at preselected bases such that codons are generated that will translate into the desired amino acids.

Another means of increasing the number of carbohydrate moieties on the LA polypeptide is by chemical or enzymatic coupling of glycosides to the polypeptide. Such methods are described in the art, e.g., in WO 87/05330 published 11 September 1987, and in Aplin and Wriston, *LA Crit. Rev. Biochem.*, pp. 259-306 (1981).

5 Removal of carbohydrate moieties present on the LA polypeptide may be accomplished chemically or enzymatically or by mutational substitution of codons encoding for amino acid residues that serve as targets for glycosylation. Chemical deglycosylation techniques are known in the art and described, for instance, by Hakimuddin, et al., *Arch. Biochem. Biophys.*, 259:52 (1987) and by Edge et al., *Anal. Biochem.*, 118:131 (1981). Enzymatic cleavage of carbohydrate moieties on polypeptides can be achieved by the use of a variety of endo-and exo-glycosidases as described by Thotakura et al., *Meth. Enzymol.*, 138:350 (1987).

10 Another type of covalent modification of LA comprises linking the LA polypeptide to one of a variety of nonproteinaceous polymers, e.g., polyethylene glycol, polypropylene glycol, or polyoxyalkylenes, in the manner set forth in U.S. Patent Nos. 4,640,835; 4,496,689; 4,301,144; 4,670,417; 4,791,192 or 15 4,179,337.

20 LA polypeptides of the present invention may also be modified in a way to form chimeric molecules comprising an LA polypeptide fused to another, heterologous polypeptide or amino acid sequence. In one embodiment, such a chimeric molecule comprises a fusion of an LA polypeptide with a tag polypeptide which provides an epitope to which an anti-tag antibody can selectively bind. The epitope tag is generally placed at the amino-or carboxyl-terminus of the LA polypeptide, although internal fusions may also be tolerated in some instances. The presence of such epitope-tagged forms of an LA polypeptide can be detected using an antibody against the tag polypeptide. Also, provision of the epitope tag enables the LA polypeptide to be readily purified by affinity purification using an anti-tag antibody or another type of affinity matrix that binds to the epitope tag. In an alternative embodiment, 25 the chimeric molecule may comprise a fusion of an LA polypeptide with an immunoglobulin or a particular region of an immunoglobulin. For a bivalent form of the chimeric molecule, such a fusion could be to the Fc region of an IgG molecule.

30 Various tag polypeptides and their respective antibodies are well known in the art. Examples include poly-histidine (poly-his) or poly-histidine-glycine (poly-his-gly) tags; the flu HA tag polypeptide and its antibody 12CA5 [Field et al., *Mol. Cell. Biol.*, 8:2159-2165 (1988)]; the c-myc tag and the 8F9, 3C7, 6E10, G4, B7 and 9E10 antibodies thereto [Evan et al., *Molecular and Cellular Biology*, 5:3610-3616 (1985)]; and the Herpes Simplex virus glycoprotein D (gD) tag and its antibody [Paborsky et al., *Protein Engineering*, 3(6):547-553 (1990)]. Other tag polypeptides include the Flag-peptide [Hopp et al., *BioTechnology*, 6:1204-1210 (1988)]; the KT3 epitope peptide [Martin et al., *Science*, 255:192-194 (1992)]; tubulin epitope peptide [Skinner et al., *J. Biol. Chem.*, 266:15163-15166 (1991)]; and the T7 gene 10 protein peptide tag [Lutz-Freyermuth et al., *Proc. Natl. Acad. Sci. USA*, 87:6393-6397 (1990)].

Also included with the definition of LA protein in one embodiment are other LA proteins of the LA family, and LA proteins from other organisms, which are cloned and expressed as outlined below. Thus, probe or degenerate polymerase chain reaction (PCR) primer sequences may be used to find other related LA proteins from humans or other organisms. As will be appreciated by those in the art, particularly useful probe and/or PCR primer sequences include the unique areas of the LA nucleic acid sequence. As is generally known in the art, preferred PCR primers are from about 15 to about 35 nucleotides in length, with from about 20 to about 30 being preferred, and may contain inosine as needed. The conditions for the PCR reaction are well known in the art.

In addition, as is outlined herein, LA proteins can be made that are longer than those encoded by the nucleic acids of the figures, for example, by the elucidation of additional sequences, the addition of epitope or purification tags, the addition of other fusion sequences, etc.

LA proteins may also be identified as being encoded by LA nucleic acids. Thus, LA proteins are encoded by nucleic acids that will hybridize to the sequences of the sequence listings, or their complements, as outlined herein.

In one embodiment, the present invention provides an LA protein referred to herein as Pik3r1 which comprises the amino acid sequence set forth in SEQ ID NO:179 and at Genbank accession number AAC52847, and which is encoded by the nucleic acid sequence set forth by nucleotides 575-2749 in SEQ ID NO:178 and at Genbank accession number U50413.

In one embodiment, the present invention provides an LA protein referred to herein as Pik3r1 which comprises the amino acid sequence set forth in SEQ ID NO:181 and at Genbank accession number A38748. In one embodiment, the present invention provides an LA protein referred to herein as Pik3r1 which is encoded by the nucleic acid sequence set forth by nucleotides 43-2217 in SEQ ID NO:180 and at Genbank accession number M61906.

In one embodiment, the present invention provides an Pik3r1 protein encoded by a nucleic acid which hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank accession number U50413.

In one embodiment, the present invention provides an Pik3r1 protein encoded by a nucleic acid which hybridizes under high stringency conditions to a nucleic acid comprising the nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank accession number M61906.

In one embodiment, the present invention provides an Pik3r1 protein encoded by a nucleic acid which comprises a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:178 and at Genbank accession number U50413.

In one embodiment, the present invention provides an Pik3r1 protein encoded by a nucleic acid which comprises a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth in SEQ ID NO:180 and at Genbank accession number M61906.

5 In one embodiment, the present invention provides an Pik3r1 protein encoded by a nucleic acid which comprises a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 575-2749 in SEQ ID NO:178 and at Genbank accession number U50413, . . .

In one embodiment, the present invention provides an Pik3r1 protein encoded by a nucleic acid which comprises a nucleic acid sequence having at least about 90% identity to the nucleic acid sequence set forth by nucleotides 43-2217 in SEQ ID NO:180 and at Genbank accession number M61906.

10 In one embodiment, the present invention provides an Pik3r1 protein comprising an SH2 domain encoded by the nucleic acid sequence set forth by nucleotides 1568-1811, or 1571-1796, or 2444-2681, or 2444-2666 in SEQ ID NO:178 and at Genbank Accession Number U50413.

15 In one embodiment, the present invention provides an Pik3r1 protein comprising an SH2 domain encoded by the nucleic acid sequence set forth by nucleotides 1037-1280, or 1040-1265, or 1913-2150, or 1913-3035 in SEQ ID NO:180 and at Genbank Accession Number M61906.

In one embodiment, the present invention provides an Pik3r1 protein comprising an SH3 domain encoded by the nucleic acid sequence set forth by nucleotides 584-797 or 593-803 in SEQ ID NO:178 and at Genbank Accession Number U50413.

20 In one embodiment, the present invention provides an Pik3r1 protein comprising an SH3 domain encoded by the nucleic acid sequence set forth by nucleotides 53-266 or 62-272 in SEQ ID NO:180 and at Genbank Accession Number M61906.

In one embodiment, the present invention provides an Pik3r1 protein comprising a RhoGAP domain encoded by the nucleic acid sequence set forth by nucleotides 998-1403 or 1001-1451 in SEQ ID NO:178 and at Genbank Accession Number U50413.

25 In one embodiment, the present invention provides an Pik3r1 protein comprising a RhoGAP domain encoded by the nucleic acid sequence set forth by nucleotides 428-929 or 428-872 in SEQ ID NO:180 and at Genbank Accession Number M61906.

In one embodiment, the present invention provides an Pik3r1 protein comprising the amino acid sequence set forth in SEQ ID NO:179 and at Genbank Accession number AAC52847.

30 In one embodiment, the present invention provides an Pik3r1 protein comprising the amino acid sequence set forth in SEQ ID NO:181 and at Genbank Accession number A38748.

In one embodiment, the present invention provides an Pik3r1 protein comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:179 and at Genbank Accession Number AAC52847.

5 In one embodiment, the present invention provides an Pik3r1 protein comprising an amino acid sequence having at least about 90% identity to the amino acid sequence set forth in SEQ ID NO:181 and at Genbank Accession Number A38748.

In one embodiment, the present invention provides an Pik3r1 protein comprising an SH2 domain comprising the amino acid sequence set forth by amino acids 332-413, or 333-408, or 624-703, or 624-698 in SEQ ID NO:179 and at Genbank Accession Number AAC52847.

10 In one embodiment, the present invention provides an Pik3r1 protein comprising an SH2 domain comprising the amino acid sequence set forth by amino acids 332-413, or 333-408, or 624-703, or 624-698 in SEQ ID NO:181 and at Genbank Accession Number A38748.

15 In one embodiment, the present invention provides an Pik3r1 protein comprising an SH3 domain comprising the amino acid sequence set forth by amino acids 4-75 or 7-77 in SEQ ID NO:179 and at Genbank Accession Number AAC52847.

In one embodiment, the present invention provides an Pik3r1 protein comprising an SH3 domain comprising the amino acid sequence set forth by amino acids 4-75 or 7-77 in SEQ ID NO:181 and at Genbank Accession Number A38748.

20 In one embodiment, the present invention provides an Pik3r1 protein comprising a RhoGAP domain comprising the amino acid sequence set forth by amino acids 142-277 or 143-293 in SEQ ID NO:179 and at Genbank Accession Number AAC52847.

In one embodiment, the present invention provides an Pik3r1 protein comprising a RhoGAP domain comprising the amino acid sequence set forth by amino acids 129-296 or 129-277 in SEQ ID NO:181 and at Genbank Accession Number A38748.

25 In a preferred embodiment, a Pik3r1 protein is a subunit of a PI3K enzyme. In a preferred embodiment, such a subunit modulates the activity of a PI3K catalytic subunit, preferably p110 as described herein. In a preferred embodiment, a Pik3r1 protein binds to phosphorylated tyrosine residues in receptor tyrosine kinases, as in the erythropoietin receptor, preferably by an SH2 domain, and tethers a PI3K catalytic subunit to the receptor. In a preferred embodiment, a Pik3r1 protein additionally binds to intracellular proteins involved in signal transduction through an SH3 domain.

30 In a preferred embodiment, a Pik3r1 protein modulates the production of phosphorylated phosphatidyl inositol lipids. In a preferred embodiment, such modulation in turn modulates the activity of

serine/threonine protein kinases, preferably PKB or PKC. In a preferred embodiment, a PIk3r1 protein modulates the phosphorylation of proteins mediating cell death and/or survival.

In a preferred embodiment, the invention provides LA antibodies. In a preferred embodiment, when the LA protein is to be used to generate antibodies, for example for immunotherapy, the LA protein should share at least one epitope or determinant with the full length protein. By "epitope" or "determinant" herein is meant a portion of a protein which will generate and/or bind an antibody or T-cell receptor in the context of MHC. Thus, in most instances, antibodies made to a smaller LA protein will be able to bind to the full length protein. In a preferred embodiment, the epitope is unique; that is, antibodies generated to a unique epitope show little or no cross-reactivity.

10 In one embodiment, the term "antibody" includes antibody fragments, as are known in the art, including Fab, Fab₂, single chain antibodies (Fv for example), chimeric antibodies, etc., either produced by the modification of whole antibodies or those synthesized de novo using recombinant DNA technologies.

15 Methods of preparing polyclonal antibodies are known to the skilled artisan. Polyclonal antibodies can be raised in a mammal, for example, by one or more injections of an immunizing agent and, if desired, an adjuvant. Typically, the immunizing agent and/or adjuvant will be injected in the mammal by multiple subcutaneous or intraperitoneal injections. The immunizing agent may include a protein encoded by a nucleic acid of the figures or fragment thereof or a fusion protein thereof. It may be useful to conjugate the immunizing agent to a protein known to be immunogenic in the mammal being immunized. Examples of such immunogenic proteins include but are not limited to keyhole limpet hemocyanin, serum albumin, bovine thyroglobulin, and soybean trypsin inhibitor. Examples of adjuvants which may be employed include Freund's complete adjuvant and MPL-TDM adjuvant (monophosphoryl Lipid A, synthetic trehalose dicorynomycolate). The immunization protocol may be selected by one skilled in the art without undue experimentation.

25 The antibodies may, alternatively, be monoclonal antibodies. Monoclonal antibodies may be prepared using hybridoma methods, such as those described by Kohler and Milstein, *Nature*, 256:495 (1975). In a hybridoma method, a mouse, hamster, or other appropriate host animal, is typically immunized with an immunizing agent to elicit lymphocytes that produce or are capable of producing antibodies that will specifically bind to the immunizing agent. Alternatively, the lymphocytes may be immunized *in vitro*. The immunizing agent will typically include a polypeptide encoded by a nucleic acid of Tables 1, 2, and 3 or fragment thereof or a fusion protein thereof. Generally, either peripheral blood lymphocytes ("PBLs") are used if cells of human origin are desired, or spleen cells or lymph node cells are used if non-human mammalian sources are desired. The lymphocytes are then fused with an immortalized cell line using a suitable fusing agent, such as polyethylene glycol, to form a hybridoma cell [Goding, *Monoclonal Antibodies: Principles and Practice*, Academic Press, (1986) pp. 59-103]. Immortalized cell lines are usually transformed mammalian cells, particularly myeloma cells of rodent, bovine and human origin. Usually, rat or mouse myeloma cell lines are employed. The hybridoma

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cells may be cultured in a suitable culture medium that preferably contains one or more substances that inhibit the growth or survival of the unfused, immortalized cells. For example, if the parental cells lack the enzyme hypoxanthine guanine phosphoribosyl transferase (HGPRT or HPRT), the culture medium for the hybridomas typically will include hypoxanthine, aminopterin, and thymidine ("HAT medium"), which substances prevent the growth of HGPRT-deficient cells.

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In one embodiment, the antibodies are bispecific antibodies. Bispecific antibodies are monoclonal, preferably human or humanized, antibodies that have binding specificities for at least two different antigens. In the present case, one of the binding specificities is for a protein encoded by a nucleic acid of the Tables 1, 2, 4, 6, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 22, 23, 24, 27, 28 or 30 or a fragment thereof, the other one is for any other antigen, and preferably for a cell-surface protein or receptor or receptor subunit, preferably one that is tumor specific.

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In a preferred embodiment, the antibodies to LA are capable of reducing or eliminating the biological function of LA, as is described below. That is, the addition of anti-LA antibodies (either polyclonal or preferably monoclonal) to LA (or cells containing LA) may reduce or eliminate the LA activity.

Generally, at least a 25% decrease in activity is preferred, with at least about 50% being particularly preferred and about a 95-100% decrease being especially preferred.

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In a preferred embodiment the antibodies to the LA proteins are humanized antibodies. Humanized forms of non-human (e.g., murine) antibodies are chimeric molecules of immunoglobulins, immunoglobulin chains or fragments thereof (such as Fv, Fab, Fab', F(ab')₂ or other antigen binding subsequences of antibodies) which contain minimal sequence derived from non-human immunoglobulin. Humanized antibodies include human immunoglobulins (recipient antibody) in which residues form a complementary determining region (CDR) of the recipient are replaced by residues from a CDR of a non-human species (donor antibody) such as mouse, rat or rabbit having the desired specificity, affinity and capacity. In some instances, Fv framework residues of the human immunoglobulin are replaced by corresponding non-human residues. Humanized antibodies may also comprise residues which are found neither in the recipient antibody nor in the imported CDR or framework sequences. In general, the humanized antibody will comprise substantially all of at least one, and typically two, variable domains, in which all or substantially all of the CDR regions correspond to those of a non-human immunoglobulin and all or substantially all of the framework residues (FR) regions are those of a human immunoglobulin consensus sequence. The humanized antibody optimally also will comprise at least a portion of an immunoglobulin constant region (Fc), typically that of a human immunoglobulin [Jones et al., Nature, 321:522-525 (1986); Riechmann et al., Nature, 332:323-329 (1988); and Presta, Curr. Op. Struct. Biol., 2:593-596 (1992)].

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Methods for humanizing non-human antibodies are well known in the art. Generally, a humanized antibody has one or more amino acid residues introduced into it from a source which is non-human. These non-human amino acid residues are often referred to as import residues, which are typically taken from an import variable domain. Humanization can be essentially performed following the

method of Winter and co-workers [Jones et al., *Nature*, 321:522-525 (1986); Riechmann et al., *Nature*, 332:323-327 (1988); Verhoeyen et al., *Science*, 239:1534-1536 (1988)], by substituting rodent CDRs or CDR sequences for the corresponding sequences of a human antibody. Accordingly, such humanized antibodies are chimeric antibodies (U.S. Patent No. 4,816,567), wherein substantially less than an intact human variable domain has been substituted by the corresponding sequence from a non-human species. In practice, humanized antibodies are typically human antibodies in which some CDR residues and possibly some FR residues are substituted by residues from analogous sites in rodent antibodies.

Human antibodies can also be produced using various techniques known in the art, including phage display libraries [Hoogenboom and Winter, *J. Mol. Biol.*, 227:381 (1991); Marks et al., *J. Mol. Biol.*, 222:581 (1991)]. The techniques of Cole et al. and Boerner et al. are also available for the preparation of human monoclonal antibodies [Cole et al., *Monoclonal Antibodies and Cancer Therapy*, Alan R. Liss, p. 77 (1985) and Boerner et al., *J. Immunol.*, 147(1):86-95 (1991)]. Similarly, human antibodies can be made by introducing human immunoglobulin loci into transgenic animals, e.g., mice in which the endogenous immunoglobulin genes have been partially or completely inactivated. Upon challenge, human antibody production is observed, which closely resembles that seen in humans in all respects, including gene rearrangement, assembly, and antibody repertoire. This approach is described, for example, in U.S. Patent Nos. 5,545,807; 5,545,806; 5,569,825; 5,625,126; 5,633,425; 5,661,016, and in the following scientific publications: Marks et al., *Bio/Technology* 10, 779-783 (1992); Lonberg et al., *Nature* 368 856-859 (1994); Morrison, *Nature* 368, 812-13 (1994); Fishwild et al., *Nature Biotechnology* 14, 845-51 (1996); Neuberger, *Nature Biotechnology* 14, 826 (1996); Lonberg and Huszar, *Intern. Rev. Immunol.* 13 65-93 (1995).

By immunotherapy is meant treatment of lymphoma with an antibody raised against an LA protein. As used herein, immunotherapy can be passive or active. Passive immunotherapy as defined herein is the passive transfer of antibody to a recipient (patient). Active immunization is the induction of antibody and/or T-cell responses in a recipient (patient). Induction of an immune response is the result of providing the recipient with an antigen to which antibodies are raised. As appreciated by one of ordinary skill in the art, the antigen may be provided by injecting a polypeptide against which antibodies are desired to be raised into a recipient, or contacting the recipient with a nucleic acid capable of expressing the antigen and under conditions for expression of the antigen.

In a preferred embodiment, oncogenes which encode secreted growth factors may be inhibited by raising antibodies against LA proteins that are secreted proteins as described above. Without being bound by theory, antibodies used for treatment, bind and prevent the secreted protein from binding to its receptor, thereby inactivating the secreted LA protein.

In a preferred embodiment, subunits of kinase holoenzymes, which holoenzymes phosphorylate substrates, preferably lipid substrates, preferably phosphatidyl inositol-conjugated lipid substrates, are

inhibited by antibodies raised against Pi3r1 proteins or portions thereof. In a preferred embodiment, such anti Pi3kr1 antibodies modulate the activity of PI3 kinase. It is recognized herein that other means of holoenzyme inhibition, preferably PI3 kinase inhibition, are known to exist and include fungal toxins, preferably wortmannin, and synthetic inhibitors, preferably LY294002.

- 5 In one embodiment, an anti-Pi3r1 antibody binds to an SH3 domain of a Pi3kr1 protein. In a preferred embodiment, such an SH3 domain comprises the amino acid sequence set forth by amino acids 4-75 or 7-77 in SEQ ID NO:179 and at Genbank accession number AAC52847. In another preferred embodiment, such an SH3 domain comprises the amino acid sequence set forth by amino acids 4-75 or 7-77 in SEQ ID NO:181 and at Genbank accession number A38748. In another 10 preferred embodiment, such an SH3 domain comprises an amino acid sequence having at least about 90% identity to the amino acid sequence set forth by amino acids 4-75 or 7-77 in SEQ ID NO:179 and at Genbank accession number AAC52847. In another preferred embodiment, such an SH3 domain comprises an amino acid sequence having at least about 90% identity to the amino acid sequence set forth by amino acids 4-75 or 7-77 in SEQ ID NO:181 and at Genbank accession number A38748.
- 15 In a preferred embodiment, an antibody recognizing an SH3 domain in a Pi3r1 protein alters the activity of Pi3r1. In a preferred embodiment, such an alteration in activity is a decrease in activity. In a preferred embodiment, such an alteration in activity alters PI3K activity. In a preferred embodiment, such an alteration in activity decreases PI3K activity.
- 20 In a preferred embodiment, an antibody recognizing an SH3 domain in a Pi3r1 protein inhibits the ability of Pi3r1 to bind to a proline rich amino acid sequence, preferably in the context of the amino acid sequence of an intracellular protein, preferably an intracellular protein involved in intracellular signal transduction.
- 25 In one embodiment, an anti-Pi3r1 antibody binds to an SH2 domain of a Pi3r1 protein. In a preferred embodiment, such an SH2 domain comprises the amino acid sequence set forth by amino acids 332-413, or 333-408, or 624-703, or 624-698 in SEQ ID NO:179 and at Genbank accession number AAC52847. In another preferred embodiment, such an SH2 domain comprises the amino acid sequence set forth by amino acids 332-413, or 333-408, or 624-703, or 624-698 in SEQ ID NO:181 and at Genbank accession number A38748. In another preferred embodiment, such an SH2 domain comprises an amino acid sequence having at least about 90% identity to the amino acid sequence set forth by amino acids 332-413, or 333-408, or 624-703, or 624-698 in SEQ ID NO:179 and at Genbank accession number AAC52847. In another preferred embodiment, such an SH2 domain comprises an amino acid sequence having at least about 90% identity to the amino acid sequence set forth by amino acids 332-413, or 333-408, or 624-703, or 624-698 in SEQ ID NO:181 and at Genbank accession number A38748.
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In a preferred embodiment, an antibody recognizing an SH2 domain in a Pik3r1 protein alters the activity of Pik3r1. In a preferred embodiment, such an alteration in activity is a decrease in activity. In a preferred embodiment, such an alteration in activity leads to a decrease in PI3K activity.

5 In a preferred embodiment, an antibody recognizing an SH2 domain in a Pik3r1 protein inhibits the ability of Pik3r1 to bind to phosphorylated tyrosine, preferably in the context of the amino acid sequence of a receptor tyrosine kinase.

In one embodiment, an anti-Pik3r1 antibody binds to a RhoGAP domain of a Pik3r1 protein. In a preferred embodiment, such a RhoGAP domain comprises the amino acid sequence set forth by amino acids 142-277 or 143-293 in SEQ ID NO:179 and at Genbank accession number AAC52847.

10 In another preferred embodiment, such a RhoGAP domain comprises the amino acid sequence set forth by amino acids 129-296 or 129-277 in SEQ ID NO:181 and at Genbank accession number A38748. In another preferred embodiment, such a RhoGAP domain comprises an amino acid sequence having at least about 90% identity to the amino acid sequence set forth by amino acids 142-277 or 143-293 in SEQ ID NO:179 and at Genbank accession number AAC52847. In another 15 preferred embodiment, such a RhoGAP domain comprises an amino acid sequence having at least about 90% identity to the amino acid sequence set forth by amino acids 129-296 or 129-277 in SEQ ID NO:181 and at Genbank accession number A38748.

20 In a preferred embodiment, an antibody recognizing a RhoGAP domain in a Pik3r1 protein alters the activity of Pik3r1. In a preferred embodiment, such an alteration in activity is a decrease in activity. In a preferred embodiment, such an alteration in activity leads to a decrease in PI3K activity.

In another preferred embodiment, the LA protein to which antibodies are raised is a transmembrane protein. Without being bound by theory, antibodies used for treatment, bind the extracellular domain of the LA protein and prevent it from binding to other proteins, such as circulating ligands or cell-associated molecules. The antibody may cause down-regulation of the transmembrane LA protein. 25 As will be appreciated by one of ordinary skill in the art, the antibody may be a competitive, non-competitive or uncompetitive inhibitor of protein binding to the extracellular domain of the LA protein. The antibody is also an antagonist of the LA protein. Further, the antibody prevents activation of the transmembrane LA protein. In one aspect, when the antibody prevents the binding of other molecules to the LA protein, the antibody prevents growth of the cell. The antibody may also sensitize the cell to 30 cytotoxic agents, including, but not limited to TNF- α , TNF- β , IL-1, INF- γ and IL-2, or chemotherapeutic agents including 5FU, vinblastine, actinomycin D, cisplatin, methotrexate, and the like. In some instances the antibody belongs to a sub-type that activates serum complement when complexed with the transmembrane protein thereby mediating cytotoxicity. Thus, lymphoma may be treated by administering to a patient antibodies directed against the transmembrane LA protein.

In another preferred embodiment, the antibody is conjugated to a therapeutic moiety. In one aspect the therapeutic moiety is a small molecule that modulates the activity of the LA protein. In another aspect the therapeutic moiety modulates the activity of molecules associated with or in close proximity to the LA protein. The therapeutic moiety may inhibit enzymatic activity such as protease or protein kinase activity associated with lymphoma.

5 In a preferred embodiment, the therapeutic moiety may also be a cytotoxic agent. In this method, targeting the cytotoxic agent to tumor tissue or cells, results in a reduction in the number of afflicted cells, thereby reducing symptoms associated with lymphoma. Cytotoxic agents are numerous and varied and include, but are not limited to, cytotoxic drugs or toxins or active fragments of such toxins.

10 Suitable toxins and their corresponding fragments include diphtheria A chain, exotoxin A chain, ricin A chain, abrin A chain, curcin, crotin, phenomycin, enomycin and the like. Cytotoxic agents also include radiochemicals made by conjugating radioisotopes to antibodies raised against LA proteins, or binding of a radionuclide to a chelating agent that has been covalently attached to the antibody. Targeting the therapeutic moiety to transmembrane LA proteins not only serves to increase the local concentration

15 of therapeutic moiety in the lymphoma, but also serves to reduce deleterious side effects that may be associated with the therapeutic moiety.

In another preferred embodiment, the LA protein against which the antibodies are raised is an intracellular protein. In this case, the antibody may be conjugated to a protein which facilitates entry into the cell. In one case, the antibody enters the cell by endocytosis. In another embodiment, a nucleic acid encoding the antibody is administered to the individual or cell. Moreover, wherein the LA protein can be targeted within a cell, i.e., the nucleus, an antibody thereto contains a signal for that target localization, i.e., a nuclear localization signal.

The LA antibodies of the invention specifically bind to LA proteins. By "specifically bind" herein is meant that the antibodies bind to the protein with a binding constant in the range of at least 10^4 - 10^6 M⁻¹, with a preferred range being 10^7 - 10^9 M⁻¹.

25 In a preferred embodiment, the LA protein is purified or isolated after expression. LA proteins may be isolated or purified in a variety of ways known to those skilled in the art depending on what other components are present in the sample. Standard purification methods include electrophoretic, molecular, immunological and chromatographic techniques, including ion exchange, hydrophobic, affinity, and reverse-phase HPLC chromatography, and chromatofocusing. For example, the LA protein may be purified using a standard anti-LA antibody column. Ultrafiltration and diafiltration techniques, in conjunction with protein concentration, are also useful. For general guidance in suitable purification techniques, see Scopes, R., Protein Purification, Springer-Verlag, NY (1982). The degree of purification necessary will vary depending on the use of the LA protein. In some instances no purification will be necessary.

Once expressed and purified if necessary, the LA proteins and nucleic acids are useful in a number of applications.

In one aspect, the expression levels of genes are determined for different cellular states in the lymphoma phenotype; that is, the expression levels of genes in normal tissue and in lymphoma tissue
5 (and in some cases, for varying severities of lymphoma that relate to prognosis, as outlined below) are evaluated to provide expression profiles. An expression profile of a particular cell state or point of development is essentially a "fingerprint" of the state; while two states may have any particular gene similarly expressed, the evaluation of a number of genes simultaneously allows the generation of a gene expression profile that is unique to the state of the cell. By comparing expression profiles of cells
10 in different states, information regarding which genes are important (including both up- and down-regulation of genes) in each of these states is obtained. Then, diagnosis may be done or confirmed: does tissue from a particular patient have the gene expression profile of normal or lymphoma tissue.

"Differential expression," or grammatical equivalents as used herein, refers to both qualitative as well as quantitative differences in the genes' temporal and/or cellular expression patterns within and
15 among the cells. Thus, a differentially expressed gene can qualitatively have its expression altered, including an activation or inactivation, in, for example, normal versus lymphoma tissue. That is, genes may be turned on or turned off in a particular state, relative to another state. As is apparent to the skilled artisan, any comparison of two or more states can be made. Such a qualitatively regulated
20 gene will exhibit an expression pattern within a state or cell type which is detectable by standard techniques in one such state or cell type, but is not detectable in both. Alternatively, the determination is quantitative in that expression is increased or decreased; that is, the expression of the gene is either upregulated, resulting in an increased amount of transcript, or downregulated, resulting in a decreased amount of transcript. The degree to which expression differs need only be large enough to quantify
25 via standard characterization techniques as outlined below, such as by use of Affymetrix GeneChip™ expression arrays, Lockhart, Nature Biotechnology, 14:1675-1680 (1996), hereby expressly incorporated by reference. Other techniques include, but are not limited to, quantitative reverse transcriptase PCR, Northern analysis and RNase protection. As outlined above, preferably the change
30 in expression (i.e. upregulation or downregulation) is at least about 50%, more preferably at least about 100%, more preferably at least about 150%, more preferably, at least about 200%, with from 300 to at least 1000% being especially preferred.

As will be appreciated by those in the art, this may be done by evaluation at either the gene transcript, or the protein level; that is, the amount of gene expression may be monitored using nucleic acid probes to the DNA or RNA equivalent of the gene transcript, and the quantification of gene expression levels, or, alternatively, the final gene product itself (protein) can be monitored, for example through
35 the use of antibodies to the LA protein and standard immunoassays (ELISAs, etc.) or other techniques, including mass spectroscopy assays, 2D gel electrophoresis assays, etc. Thus, the proteins corresponding to LA genes, i.e. those identified as being important in a lymphoma phenotype, can be evaluated in a lymphoma diagnostic test.

In a preferred embodiment, gene expression monitoring is done and a number of genes, i.e. an expression profile, is monitored simultaneously, although multiple protein expression monitoring can be done as well. Similarly, these assays may be done on an individual basis as well.

In this embodiment, the LA nucleic acid probes may be attached to biochips as outlined herein for the detection and quantification of LA sequences in a particular cell. The assays are done as is known in the art. As will be appreciated by those in the art, any number of different LA sequences may be used as probes, with single sequence assays being used in some cases, and a plurality of the sequences described herein being used in other embodiments. In addition, while solid-phase assays are described, any number of solution based assays may be done as well.

In a preferred embodiment, both solid and solution based assays may be used to detect LA sequences that are up-regulated or down-regulated in lymphoma as compared to normal lymphoid tissue. In instances where the LA sequence has been altered but shows the same expression profile or an altered expression profile, the protein will be detected as outlined herein.

In a preferred embodiment nucleic acids encoding the LA protein are detected. Although DNA or RNA encoding the LA protein may be detected, of particular interest are methods wherein the mRNA encoding a LA protein is detected. The presence of mRNA in a sample is an indication that the LA gene has been transcribed to form the mRNA, and suggests that the protein is expressed. Probes to detect the mRNA can be any nucleotide/deoxynucleotide probe that is complementary to and base pairs with the mRNA and includes but is not limited to oligonucleotides, cDNA or RNA. Probes also should contain a detectable label, as defined herein. In one method the mRNA is detected after immobilizing the nucleic acid to be examined on a solid support such as nylon membranes and hybridizing the probe with the sample. Following washing to remove the non-specifically bound probe, the label is detected. In another method detection of the mRNA is performed *in situ*. In this method permeabilized cells or tissue samples are contacted with a detectably labeled nucleic acid probe for sufficient time to allow the probe to hybridize with the target mRNA. Following washing to remove the non-specifically bound probe, the label is detected. For example a digoxygenin labeled riboprobe (RNA probe) that is complementary to the mRNA encoding a LA protein is detected by binding the digoxygenin with an anti-digoxygenin secondary antibody and developed with nitro blue tetrazolium and 5-bromo-4-chloro-3-indoyl phosphate.

In a preferred embodiment, any of the three classes of proteins as described herein (secreted, transmembrane or intracellular proteins) are used in diagnostic assays. The LA proteins, antibodies, nucleic acids, modified proteins and cells containing LA sequences are used in diagnostic assays. This can be done on an individual gene or corresponding polypeptide level, or as sets of assays.

As described and defined herein, LA proteins find use as markers of lymphoma. Detection of these proteins in putative lymphomeric tissue or patients allows for a determination or diagnosis of lymphoma. Numerous methods known to those of ordinary skill in the art find use in detecting lymphoma. In one

embodiment, antibodies are used to detect LA proteins. A preferred method separates proteins from a sample or patient by electrophoresis on a gel (typically a denaturing and reducing protein gel, but may be any other type of gel including isoelectric focusing gels and the like). Following separation of proteins, the LA protein is detected by immunoblotting with antibodies raised against the LA protein.

5 Methods of immunoblotting are well known to those of ordinary skill in the art.

In another preferred method, antibodies to the LA protein find use in *in situ* imaging techniques. In this method cells are contacted with from one to many antibodies to the LA protein(s). Following washing to remove non-specific antibody binding, the presence of the antibody or antibodies is detected. In one embodiment the antibody is detected by incubating with a secondary antibody that contains a detectable label. In another method the primary antibody to the LA protein(s) contains a detectable label. In another preferred embodiment each one of multiple primary antibodies contains a distinct and detectable label. This method finds particular use in simultaneous screening for a plurality of LA proteins. As will be appreciated by one of ordinary skill in the art, numerous other histological imaging techniques are useful in the invention.

15 In a preferred embodiment the label is detected in a fluorometer which has the ability to detect and distinguish emissions of different wavelengths. In addition, a fluorescence activated cell sorter (FACS) can be used in the method.

In another preferred embodiment, antibodies find use in diagnosing lymphoma from blood samples. As previously described, certain LA proteins are secreted/circulating molecules. Blood samples, therefore, are useful as samples to be probed or tested for the presence of secreted LA proteins. 20 Antibodies can be used to detect the LA by any of the previously described immunoassay techniques including ELISA, immunoblotting (Western blotting), immunoprecipitation, BIACORE technology and the like, as will be appreciated by one of ordinary skill in the art.

25 In a preferred embodiment, *in situ* hybridization of labeled LA nucleic acid probes to tissue arrays is done. For example, arrays of tissue samples, including LA tissue and/or normal tissue, are made. *In situ* hybridization as is known in the art can then be done.

It is understood that when comparing the expression fingerprints between an individual and a standard, the skilled artisan can make a diagnosis as well as a prognosis. It is further understood that the genes which indicate the diagnosis may differ from those which indicate the prognosis.

30 In a preferred embodiment, the LA proteins, antibodies, nucleic acids, modified proteins and cells containing LA sequences are used in prognosis assays. As above, gene expression profiles can be generated that correlate to lymphoma severity, in terms of long term prognosis. Again, this may be done on either a protein or gene level, with the use of genes being preferred. As above, the LA

probes are attached to biochips for the detection and quantification of LA sequences in a tissue or patient. The assays proceed as outlined for diagnosis.

In a preferred embodiment, any of the LA sequences as described herein are used in drug screening assays. The LA proteins, antibodies, nucleic acids, modified proteins and cells containing LA sequences are used in drug screening assays or by evaluating the effect of drug candidates on a "gene expression profile" or expression profile of polypeptides. In one embodiment, the expression profiles are used, preferably in conjunction with high throughput screening techniques to allow monitoring for expression profile genes after treatment with a candidate agent, Zlokarnik, et al., Science 279, 84-8 (1998), Heid, et al., Genome Res., 6:986-994 (1996).

In a preferred embodiment, the LA proteins, antibodies, nucleic acids, modified proteins and cells containing the native or modified LA proteins are used in screening assays. That is, the present invention provides novel methods for screening for compositions which modulate the lymphoma phenotype. As above, this can be done by screening for modulators of gene expression or for modulators of protein activity. Similarly, this may be done on an individual gene or protein level or by evaluating the effect of drug candidates on a "gene expression profile". In a preferred embodiment, the expression profiles are used, preferably in conjunction with high throughput screening techniques to allow monitoring for expression profile genes after treatment with a candidate agent, see Zlokarnik, supra.

Having identified the LA genes herein, a variety of assays to evaluate the effects of agents on gene expression may be executed. In a preferred embodiment, assays may be run on an individual gene or protein level. That is, having identified a particular gene as aberrantly regulated in lymphoma, candidate bioactive agents may be screened to modulate the gene's response. "Modulation" thus includes both an increase and a decrease in gene expression or activity. The preferred amount of modulation will depend on the original change of the gene expression in normal versus tumor tissue, with changes of at least 10%, preferably 50%, more preferably 100-300%, and in some embodiments 300-1000% or greater. Thus, if a gene exhibits a 4 fold increase in tumor compared to normal tissue, a decrease of about four fold is desired; a 10 fold decrease in tumor compared to normal tissue gives a 10 fold increase in expression for a candidate agent is desired, etc. Alternatively, where the LA sequence has been altered but shows the same expression profile or an altered expression profile, the protein will be detected as outlined herein.

As will be appreciated by those in the art, this may be done by evaluation at either the gene or the protein level; that is, the amount of gene expression may be monitored using nucleic acid probes and the quantification of gene expression levels, or, alternatively, the level of the gene product itself can be monitored, for example through the use of antibodies to the LA protein and standard immunoassays.

Alternatively, binding and bioactivity assays with the protein may be done as outlined below.

In a preferred embodiment, gene expression monitoring is done and a number of genes, i.e. an expression profile, is monitored simultaneously, although multiple protein expression monitoring can be done as well.

5 In this embodiment, the LA nucleic acid probes are attached to biochips as outlined herein for the detection and quantification of LA sequences in a particular cell. The assays are further described below.

10 Generally, in a preferred embodiment, a candidate bioactive agent is added to the cells prior to analysis. Moreover, screens are provided to identify a candidate bioactive agent which modulates lymphoma, modulates LA proteins, binds to a LA protein, or interferes between the binding of a LA protein and an antibody.

15 The term "candidate bioactive agent" or "drug candidate" or grammatical equivalents as used herein describes any molecule, e.g., protein, oligopeptide, small organic or inorganic molecule, polysaccharide, polynucleotide, etc., to be tested for bioactive agents that are capable of directly or indirectly altering either the lymphoma phenotype, binding to and/or modulating the bioactivity of an LA protein, or the expression of a LA sequence, including both nucleic acid sequences and protein sequences. In a particularly preferred embodiment, the candidate agent suppresses a LA phenotype, for example to a normal tissue fingerprint. Similarly, the candidate agent preferably suppresses a severe LA phenotype. Generally a plurality of assay mixtures are run in parallel with different agent concentrations to obtain a differential response to the various concentrations. Typically, one of these 20 concentrations serves as a negative control, i.e., at zero concentration or below the level of detection.

25 In one aspect, a candidate agent will neutralize the effect of an LA protein. By "neutralize" is meant that activity of a protein is either inhibited or counter acted against so as to have substantially no effect on a cell.

30 Candidate agents encompass numerous chemical classes, though typically they are organic or inorganic molecules, preferably small organic compounds having a molecular weight of more than 100 and less than about 2,500 daltons. Preferred small molecules are less than 2000, or less than 1500 or less than 1000 or less than 500 D. Candidate agents comprise functional groups necessary for structural interaction with proteins, particularly hydrogen bonding, and typically include at least an amine, carbonyl, hydroxyl or carboxyl group, preferably at least two of the functional chemical groups. The candidate agents often comprise cyclical carbon or heterocyclic structures and/or aromatic or polyaromatic structures substituted with one or more of the above functional groups. Candidate agents are also found among biomolecules including peptides, saccharides, fatty acids, steroids, purines, pyrimidines, derivatives, structural analogs or combinations thereof. Particularly preferred are peptides.

Candidate agents are obtained from a wide variety of sources including libraries of synthetic or natural compounds. For example, numerous means are available for random and directed synthesis of a wide variety of organic compounds and biomolecules, including expression of randomized oligonucleotides. Alternatively, libraries of natural compounds in the form of bacterial, fungal, plant and animal extracts are available or readily produced. Additionally, natural or synthetically produced libraries and compounds are readily modified through conventional chemical, physical and biochemical means. Known pharmacological agents may be subjected to directed or random chemical modifications, such as acylation, alkylation, esterification, amidification to produce structural analogs.

- 5 In a preferred embodiment, the candidate bioactive agents are proteins. By "protein" herein is meant at least two covalently attached amino acids, which includes proteins, polypeptides, oligopeptides and peptides. The protein may be made up of naturally occurring amino acids and peptide bonds, or synthetic peptidomimetic structures. Thus "amino acid", or "peptide residue", as used herein means both naturally occurring and synthetic amino acids. For example, homo-phenylalanine, citrulline and noreleucine are considered amino acids for the purposes of the invention. "Amino acid" also includes imino acid residues such as proline and hydroxyproline. The side chains may be in either the (R) or the (S) configuration. In the preferred embodiment, the amino acids are in the (S) or L-configuration. If non-naturally occurring side chains are used, non-amino acid substituents may be used, for example to prevent or retard in vivo degradations.
- 10 In a preferred embodiment, the candidate bioactive agents are naturally occurring proteins or fragments of naturally occurring proteins. Thus, for example, cellular extracts containing proteins, or random or directed digests of proteinaceous cellular extracts, may be used. In this way libraries of prokaryotic and eucaryotic proteins may be made for screening in the methods of the invention. Particularly preferred in this embodiment are libraries of bacterial, fungal, viral, and mammalian proteins, with the latter being preferred, and human proteins being especially preferred.
- 15 In a preferred embodiment, the candidate bioactive agents are peptides of from about 5 to about 30 amino acids, with from about 5 to about 20 amino acids being preferred, and from about 7 to about 15 being particularly preferred. The peptides may be digests of naturally occurring proteins as is outlined

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sequence are either held constant, or are selected from a limited number of possibilities. For example, in a preferred embodiment, the nucleotides or amino acid residues are randomized within a defined class, for example, of hydrophobic amino acids, hydrophilic residues, sterically biased (either small or large) residues, towards the creation of nucleic acid binding domains, the creation of cysteines, for cross-linking, prolines for SH-3 domains, serines, threonines, tyrosines or histidines for phosphorylation sites, etc., or to purines, etc.

In a preferred embodiment, the candidate bioactive agents are nucleic acids, as defined above.

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As described above generally for proteins, nucleic acid candidate bioactive agents may be naturally occurring nucleic acids, random nucleic acids, or "biased" random nucleic acids. For example, digests of prokaryotic or eucaryotic genomes may be used as is outlined above for proteins.

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In a preferred embodiment, the candidate bioactive agents are organic chemical moieties, a wide variety of which are available in the literature.

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In assays for altering the expression profile of one or more LA genes, after the candidate agent has been added and the cells allowed to incubate for some period of time, the sample containing the target sequences to be analyzed is added to the biochip. If required, the target sequence is prepared using known techniques. For example, the sample may be treated to lyse the cells, using known lysis buffers, electroporation, etc., with purification and/or amplification such as PCR occurring as needed, as will be appreciated by those in the art. For example, an *in vitro* transcription with labels covalently attached to the nucleosides is done. Generally, the nucleic acids are labeled with a label as defined herein, with biotin-FITC or PE, cy3 and cy5 being particularly preferred.

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In a preferred embodiment, the target sequence is labeled with, for example, a fluorescent, chemiluminescent, chemical, or radioactive signal, to provide a means of detecting the target sequence's specific binding to a probe. The label also can be an enzyme, such as, alkaline phosphatase or horseradish peroxidase, which when provided with an appropriate substrate produces a product that can be detected. Alternatively, the label can be a labeled compound or small molecule, such as an enzyme inhibitor, that binds but is not catalyzed or altered by the enzyme. The label also can be a moiety or compound, such as, an epitope tag or biotin which specifically binds to streptavidin. For the example of biotin, the streptavidin is labeled as described above, thereby, providing a detectable signal for the bound target sequence. As known in the art, unbound labeled streptavidin is removed prior to analysis.

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As will be appreciated by those in the art, these assays can be direct hybridization assays or can comprise "sandwich assays", which include the use of multiple probes, as is generally outlined in U.S. Patent Nos. 5,681,702, 5,597,909, 5,545,730, 5,594,117, 5,591,584, 5,571,670, 5,580,731, 5,571,670, 5,591,584, 5,624,802, 5,635,352, 5,594,118, 5,359,100, 5,124,246 and 5,681,697, all of which are hereby incorporated by reference. In this embodiment, in general, the target nucleic acid is prepared

as outlined above, and then added to the biochip comprising a plurality of nucleic acid probes, under conditions that allow the formation of a hybridization complex.

5 A variety of hybridization conditions may be used in the present invention, including high, moderate and low stringency conditions as outlined above. The assays are generally run under stringency conditions which allows formation of the label probe hybridization complex only in the presence of target. Stringency can be controlled by altering a step parameter that is a thermodynamic variable, including, but not limited to, temperature, formamide concentration, salt concentration, chaotropic salt concentration pH, organic solvent concentration, etc.

10 These parameters may also be used to control non-specific binding, as is generally outlined in U.S. Patent No. 5,681,697. Thus it may be desirable to perform certain steps at higher stringency conditions to reduce non-specific binding.

15 The reactions outlined herein may be accomplished in a variety of ways, as will be appreciated by those in the art. Components of the reaction may be added simultaneously, or sequentially, in any order, with preferred embodiments outlined below. In addition, the reaction may include a variety of other reagents may be included in the assays. These include reagents like salts, buffers, neutral proteins, e.g. albumin, detergents, etc which may be used to facilitate optimal hybridization and detection, and/or reduce non-specific or background interactions. Also reagents that otherwise improve the efficiency of the assay, such as protease inhibitors, nuclease inhibitors, anti-microbial agents, etc., may be used, depending on the sample preparation methods and purity of the target. In 20 addition, either solid phase or solution based (i.e., kinetic PCR) assays may be used.

Once the assay is run, the data is analyzed to determine the expression levels, and changes in expression levels as between states, of individual genes, forming a gene expression profile.

25 In a preferred embodiment, as for the diagnosis and prognosis applications, having identified the differentially expressed gene(s) or mutated gene(s) important in any one state, screens can be run to alter the expression of the genes individually. That is, screening for modulation of regulation of expression of a single gene can be done. Thus, for example, particularly in the case of target genes whose presence or absence is unique between two states, screening is done for modulators of the target gene expression.

30 In addition screens can be done for novel genes that are induced in response to a candidate agent. After identifying a candidate agent based upon its ability to suppress a LA expression pattern leading to a normal expression pattern, or modulate a single LA gene expression profile so as to mimic the expression of the gene from normal tissue, a screen as described above can be performed to identify genes that are specifically modulated in response to the agent. Comparing expression profiles between normal tissue and agent treated LA tissue reveals genes that are not expressed in normal 35 tissue or LA tissue, but are expressed in agent treated tissue. These agent specific sequences can be

identified and used by any of the methods described herein for LA genes or proteins. In particular these sequences and the proteins they encode find use in marking or identifying agent treated cells. In addition, antibodies can be raised against the agent induced proteins and used to target novel therapeutics to the treated LA tissue sample.

5 Thus, in one embodiment, a candidate agent is administered to a population of LA cells, that thus has an associated LA expression profile. By "administration" or "contacting" herein is meant that the candidate agent is added to the cells in such a manner as to allow the agent to act upon the cell, whether by uptake and intracellular action, or by action at the cell surface. In some embodiments, nucleic acid encoding a proteinaceous candidate agent (i.e. a peptide) may be put into a viral construct such as a retroviral construct and added to the cell, such that expression of the peptide agent is accomplished; see PCT US97/01019, hereby expressly incorporated by reference.

10 Once the candidate agent has been administered to the cells, the cells can be washed if desired and are allowed to incubate under preferably physiological conditions for some period of time. The cells are then harvested and a new gene expression profile is generated, as outlined herein.

15 Thus, for example, LA tissue may be screened for agents that reduce or suppress the LA phenotype. A change in at least one gene of the expression profile indicates that the agent has an effect on LA activity. By defining such a signature for the LA phenotype, screens for new drugs that alter the phenotype can be devised. With this approach, the drug target need not be known and need not be represented in the original expression screening platform, nor does the level of transcript for the target 20 protein need to change.

In a preferred embodiment, as outlined above, screens may be done on individual genes and gene products (proteins). That is, having identified a particular differentially expressed gene as important in a particular state, screening of modulators of either the expression of the gene or the gene product itself can be done. The gene products of differentially expressed genes are sometimes referred to herein as "LA proteins" or an "LAP". The LAP may be a fragment, or alternatively, be the full length protein to the fragment encoded by the nucleic acids of the figures. Preferably, the LAP is a fragment. 25 In another embodiment, the sequences are sequence variants as further described herein.

30 Preferably, the LAP is a fragment of approximately 14 to 24 amino acids long. More preferably the fragment is a soluble fragment. Preferably, the fragment includes a non-transmembrane region. In a preferred embodiment, the fragment has an N-terminal Cys to aid in solubility. In one embodiment, the c-terminus of the fragment is kept as a free acid and the n-terminus is a free amine to aid in coupling, i.e., to cysteine.

In one embodiment the LA proteins are conjugated to an immunogenic agent as discussed herein. In one embodiment the LA protein is conjugated to BSA.

In a preferred embodiment, screening is done to alter the biological function of the expression product of the LA gene. Again, having identified the importance of a gene in a particular state, screening for agents that bind and/or modulate the biological activity of the gene product can be run as is more fully outlined below.

5 In a preferred embodiment, screens are designed to first find candidate agents that can bind to LA proteins, and then these agents may be used in assays that evaluate the ability of the candidate agent to modulate the LAP activity and the lymphoma phenotype. Thus, as will be appreciated by those in the art, there are a number of different assays which may be run; binding assays and activity assays.

10 In a preferred embodiment, binding assays are done. In general, purified or isolated gene product is used; that is, the gene products of one or more LA nucleic acids are made. In general, this is done as is known in the art. For example, antibodies are generated to the protein gene products, and standard immunoassays are run to determine the amount of protein present. Alternatively, cells comprising the LA proteins can be used in the assays.

15 Thus, in a preferred embodiment, the methods comprise combining a LA protein and a candidate bioactive agent, and determining the binding of the candidate agent to the LA protein. Preferred embodiments utilize the human or mouse LA protein, although other mammalian proteins may also be used, for example for the development of animal models of human disease. In some embodiments, as outlined herein, variant or derivative LA proteins may be used.

20 Generally, in a preferred embodiment of the methods herein, the LA protein or the candidate agent is non-diffusably bound to an insoluble support having isolated sample receiving areas (e.g. a microtiter plate, an array, etc.). The insoluble supports may be made of any composition to which the compositions can be bound, is readily separated from soluble material, and is otherwise compatible with the overall method of screening. The surface of such supports may be solid or porous and of any convenient shape. Examples of suitable insoluble supports include microtiter plates, arrays, 25 membranes and beads. These are typically made of glass, plastic (e.g., polystyrene), polysaccharides, nylon or nitrocellulose, teflon™, etc. Microtiter plates and arrays are especially convenient because a large number of assays can be carried out simultaneously, using small amounts of reagents and samples. The particular manner of binding of the composition is not crucial so long as it is compatible with the reagents and overall methods of the invention, maintains the activity of the 30 composition and is nondiffusible. Preferred methods of binding include the use of antibodies (which do not sterically block either the ligand binding site or activation sequence when the protein is bound to the support), direct binding to "sticky" or ionic supports, chemical crosslinking, the synthesis of the protein or agent on the surface, etc. Following binding of the protein or agent, excess unbound material is removed by washing. The sample receiving areas may then be blocked through incubation 35 with bovine serum albumin (BSA), casein or other innocuous protein or other moiety.

In a preferred embodiment, the LA protein is bound to the support, and a candidate bioactive agent is added to the assay. Alternatively, the candidate agent is bound to the support and the LA protein is added. Novel binding agents include specific antibodies, non-natural binding agents identified in screens of chemical libraries, peptide analogs, etc. Of particular interest are screening assays for agents that have a low toxicity for human cells. A wide variety of assays may be used for this purpose, including labeled *in vitro* protein-protein binding assays, electrophoretic mobility shift assays, immunoassays for protein binding, functional assays (phosphorylation assays, etc.) and the like...

The determination of the binding of the candidate bioactive agent to the LA protein may be done in a number of ways. In a preferred embodiment, the candidate bioactive agent is labeled, and binding determined directly. For example, this may be done by attaching all or a portion of the LA protein to a solid support, adding a labeled candidate agent (for example a fluorescent label), washing off excess reagent, and determining whether the label is present on the solid support. Various blocking and washing steps may be utilized as is known in the art.

By "labeled" herein is meant that the compound is either directly or indirectly labeled with a label which provides a detectable signal, e.g. radioisotope, fluorescers, enzyme, antibodies, particles such as magnetic particles, chemiluminescers, or specific binding molecules, etc. Specific binding molecules include pairs, such as biotin and streptavidin, digoxin and antidigoxin etc. For the specific binding members, the complementary member would normally be labeled with a molecule which provides for detection, in accordance with known procedures, as outlined above. The label can directly or indirectly provide a detectable signal.

In some embodiments, only one of the components is labeled. For example, the proteins (or proteinaceous candidate agents) may be labeled at tyrosine positions using ^{125}I , or with fluorophores. Alternatively, more than one component may be labeled with different labels; using ^{125}I for the proteins, for example, and a fluorophor for the candidate agents.

In a preferred embodiment, the binding of the candidate bioactive agent is determined through the use of competitive binding assays. In this embodiment, the competitor is a binding moiety known to bind to the target molecule (i.e. LA protein), such as an antibody, peptide, binding partner, ligand, etc. Under certain circumstances, there may be competitive binding as between the bioactive agent and the binding moiety, with the binding moiety displacing the bioactive agent.

In a preferred embodiment, the Nrf2 binding moiety is a nucleic acid comprising the Nrf2 binding sequence GCTGAGTCATGATGAGTCA. In another preferred embodiment, the Nrf2 binding moiety is a transcriptional cofactor involved in Nrf2-mediated gene regulation. In a preferred embodiment, the DNA binding domain of Nrf2 is used in binding assays. In one embodiment, the transcriptional activation domain of Nrf2 is used in binding assays.

In one embodiment, the candidate bioactive agent is labeled. Either the candidate bioactive agent, or the competitor, or both, is added first to the protein for a time sufficient to allow binding, if present. Incubations may be performed at any temperature which facilitates optimal activity, typically between 4 and 40°C. Incubation periods are selected for optimum activity, but may also be optimized to facilitate rapid high through put screening. Typically between 0.1 and 1 hour will be sufficient. Excess reagent is generally removed or washed away. The second component is then added, and the presence or absence of the labeled component is followed, to indicate binding.

5 In a preferred embodiment, the competitor is added first, followed by the candidate bioactive agent. Displacement of the competitor is an indication that the candidate bioactive agent is binding to the LA protein and thus is capable of binding to, and potentially modulating, the activity of the LA protein. In 10 this embodiment, either component can be labeled. Thus, for example, if the competitor is labeled, the presence of label in the wash solution indicates displacement by the agent. Alternatively, if the candidate bioactive agent is labeled, the presence of the label on the support indicates displacement.

15 In an alternative embodiment, the candidate bioactive agent is added first, with incubation and washing, followed by the competitor. The absence of binding by the competitor may indicate that the bioactive agent is bound to the LA protein with a higher affinity. Thus, if the candidate bioactive agent is labeled, the presence of the label on the support, coupled with a lack of competitor binding, may indicate that the candidate agent is capable of binding to the LA protein.

20 In a preferred embodiment, the methods comprise differential screening to identify bioactive agents that are capable of modulating the activity of the LA proteins. In this embodiment, the methods comprise combining a LA protein and a competitor in a first sample. A second sample comprises a candidate bioactive agent, a LA protein and a competitor. The binding of the competitor is determined for both samples, and a change, or difference in binding between the two samples indicates the presence of an agent capable of binding to the LA protein and potentially modulating its activity. That 25 is, if the binding of the competitor is different in the second sample relative to the first sample, the agent is capable of binding to the LA protein.

30 Alternatively, a preferred embodiment utilizes differential screening to identify drug candidates that bind to the native LA protein, but cannot bind to modified LA proteins. The structure of the LA protein may be modeled, and used in rational drug design to synthesize agents that interact with that site. Drug candidates that affect LA bioactivity are also identified by screening drugs for the ability to either enhance or reduce the activity of the protein.

35 In a preferred embodiment, transcription assays as known in the art, for example as disclosed in (Ausubel, *supra*) and Caterina et al., NAR 22:2383-2391, 1994, are used in screens to identify candidate bioactive agents that can affect Nrf2 protein activity, particularly transcription regulating activity. In a preferred embodiment, the transcription assays employ the Nrf2 DNA binding sequence GCTGAGTCATGATGAGTCA. In a preferred embodiment, an Nrf2 protein comprises the amino acid

sequence set forth in SEQ ID NO:211 and at Genbank accession number AAA68291, or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth in SEQ ID NO:213 and at Genbank accession number NP_006155, or a fragment thereof. In another preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth by amino acids 477 to 518 in SEQ ID NO:211 and at Genbank accession number AAA68291. In another preferred embodiment, an Nrf2 protein comprises the amino acid sequence set forth by amino acids 482 to 526, more preferably 482 to 504, in SEQ ID NO:213 and at Genbank accession number NP_006155.

In one embodiment, the portion of Nrf2 protein used comprises the DNA binding domain, such as the basic domain of a basic leucine zipper domain-containing protein. In one embodiment, the portion of Nrf2 used comprises the transcriptional activation domain, such as the acidic domain of a basic leucine zipper domain-containing protein.

Positive controls and negative controls may be used in the assays. Preferably all control and test samples are performed in at least triplicate to obtain statistically significant results. Incubation of all samples is for a time sufficient for the binding of the agent to the protein. Following incubation, all samples are washed free of non-specifically bound material and the amount of bound, generally labeled agent determined. For example, where a radiolabel is employed, the samples may be counted in a scintillation counter to determine the amount of bound compound.

A variety of other reagents may be included in the screening assays. These include reagents like salts, neutral proteins, e.g. albumin, detergents, etc which may be used to facilitate optimal protein-protein binding and/or reduce non-specific or background interactions. Also reagents that otherwise improve the efficiency of the assay, such as protease inhibitors, nuclease inhibitors, anti-microbial agents, etc., may be used. The mixture of components may be added in any order that provides for the requisite binding.

Screening for agents that modulate the activity of LA proteins may also be done. In a preferred embodiment, methods for screening for a bioactive agent capable of modulating the activity of LA proteins comprise the steps of adding a candidate bioactive agent to a sample of LA proteins, as above, and determining an alteration in the biological activity of LA proteins. "Modulating the activity of an LA protein" includes an increase in activity, a decrease in activity, or a change in the type or kind of activity present. Thus, in this embodiment, the candidate agent should both bind to LA proteins (although this may not be necessary), and alter its biological or biochemical activity as defined herein. The methods include both *in vitro* screening methods, as are generally outlined above, and *in vivo* screening of cells for alterations in the presence, distribution, activity or amount of LA proteins.

Thus, in this embodiment, the methods comprise combining a LA sample and a candidate bioactive agent, and evaluating the effect on LA activity. By "LA activity" or grammatical equivalents herein is meant one of the LA protein's biological activities, including, but not limited to, its role in lymphoma,

including cell division, preferably in lymphoid tissue, cell proliferation, tumor growth and transformation of cells. In one embodiment, LA activity includes activation of or by a protein encoded by a nucleic acid of the table. An inhibitor of LA activity is the inhibition of any one or more LA activities.

5 In a preferred embodiment, the activity of the LA protein is increased; in another preferred embodiment, the activity of the LA protein is decreased. Thus, bioactive agents that are antagonists are preferred in some embodiments, and bioactive agents that are agonists may be preferred in other embodiments.

10 In a preferred embodiment, the invention provides methods for screening for bioactive agents capable of modulating the activity of a LA protein. The methods comprise adding a candidate bioactive agent, as defined above, to a cell comprising LA proteins. Preferred cell types include almost any cell. The cells contain a recombinant nucleic acid that encodes a LA protein. In a preferred embodiment, a library of candidate agents are tested on a plurality of cells.

15 In one aspect, the assays are evaluated in the presence or absence or previous or subsequent exposure of physiological signals, for example hormones, antibodies, peptides, antigens, cytokines, growth factors, action potentials, pharmacological agents including chemotherapeutics, radiation, carcinogenics, or other cells (i.e. cell-cell contacts). In another example, the determinations are determined at different stages of the cell cycle process.

20 In this way, bioactive agents are identified. Compounds with pharmacological activity are able to enhance or interfere with the activity of the LA protein.

25 In one embodiment, a method of inhibiting lymphoma cancer cell division is provided. The method comprises administration of a lymphoma cancer inhibitor.

In another embodiment, a method of inhibiting tumor growth is provided. The method comprises administration of a lymphoma cancer inhibitor.

30 In a further embodiment, methods of treating cells or individuals with cancer are provided. The method comprises administration of a lymphoma cancer inhibitor.

In one embodiment, a lymphoma cancer inhibitor is an antibody as discussed above. In another embodiment, the lymphoma cancer inhibitor is an antisense molecule. Antisense molecules as used herein include antisense or sense oligonucleotides comprising a single-stranded nucleic acid sequence (either RNA or DNA) capable of binding to target mRNA (sense) or DNA (antisense) sequences for lymphoma cancer molecules. Antisense or sense oligonucleotides, according to the present invention, comprise a fragment generally at least about 14 nucleotides, preferably from about 14 to 30 nucleotides. The ability to derive an antisense or a sense oligonucleotide, based upon a cDNA

sequence encoding a given protein is described in, for example, Stein and Cohen, *Cancer Res.* 48:2659, (1988) and van der Krol et al., *BioTechniques* 6:958, (1988).

Antisense molecules may be introduced into a cell containing the target nucleotide sequence by formation of a conjugate with a ligand binding molecule, as described in WO 91/04753. Suitable ligand binding molecules include, but are not limited to, cell surface receptors, growth factors, other cytokines, or other ligands that bind to cell surface receptors. Preferably, conjugation of the ligand binding molecule does not substantially interfere with the ability of the ligand binding molecule to bind to its corresponding molecule or receptor, or block entry of the sense or antisense oligonucleotide or its conjugated version into the cell. Alternatively, a sense or an antisense oligonucleotide may be introduced into a cell containing the target nucleic acid sequence by formation of an oligonucleotide-lipid complex, as described in WO 90/10448. It is understood that the use of antisense molecules or knock out and knock in models may also be used in screening assays as discussed above, in addition to methods of treatment.

The compounds having the desired pharmacological activity may be administered in a physiologically acceptable carrier to a host, as previously described. The agents may be administered in a variety of ways, orally, parenterally e.g., subcutaneously, intraperitoneally, intravascularly, etc. Depending upon the manner of introduction, the compounds may be formulated in a variety of ways. The concentration of therapeutically active compound in the formulation may vary from about 0.1-100% wgt/vol. The agents may be administered alone or in combination with other treatments, i.e., radiation.

The pharmaceutical compositions can be prepared in various forms, such as granules, tablets, pills, suppositories, capsules, suspensions, salves, lotions and the like. Pharmaceutical grade organic or inorganic carriers and/or diluents suitable for oral and topical use can be used to make up compositions containing the therapeutically-active compounds. Diluents known to the art include aqueous media, vegetable and animal oils and fats. Stabilizing agents, wetting and emulsifying agents, salts for varying the osmotic pressure or buffers for securing an adequate pH value, and skin penetration enhancers can be used as auxiliary agents.

Without being bound by theory, it appears that the various LA sequences are important in lymphoma. Accordingly, disorders based on mutant or variant LA genes may be determined. In one embodiment, the invention provides methods for identifying cells containing variant LA genes comprising determining all or part of the sequence of at least one endogenous LA genes in a cell. As will be appreciated by those in the art, this may be done using any number of sequencing techniques. In a preferred embodiment, the invention provides methods of identifying the LA genotype of an individual comprising determining all or part of the sequence of at least one LA gene of the individual. This is generally done in at least one tissue of the individual, and may include the evaluation of a number of tissues or different samples of the same tissue. The method may include comparing the sequence of the sequenced LA gene to a known LA gene, i.e., a wild-type gene. As will be appreciated by those in

the art, alterations in the sequence of some oncogenes can be an indication of either the presence of the disease, or propensity to develop the disease, or prognosis evaluations.

The sequence of all or part of the LA gene can then be compared to the sequence of a known LA gene to determine if any differences exist. This can be done using any number of known homology programs, such as Bestfit, etc. In a preferred embodiment, the presence of a difference in the sequence between the LA gene of the patient and the known LA gene is indicative of a disease state or a propensity for a disease state, as outlined herein.

It will be recognized that in some cases, particularly those concerning tumor suppresser genes, or recessive mutations generally, Nrf2 sequences characteristic of an Nrf2 phenotype will be found in normal lymphoid tissue. In these case it will be recognized that other Nrf2 gene alleles found in the tissue are likely involved in the maintenance of the normal lymphoid phenotype.

It will also be recognized that many transcription factors function as multimers, and as such, dominant negative effects in respect of the physiological processes they regulate are often encountered with altered alleles. That is, a single alternate allele (alternate in respect of the recognized wildtype allele) is often sufficient to alter transcription as normally regulated by wildtype protein, through protein-protein interactions and the dominant dysfunction of an alternate protein.

In a preferred embodiment, the LA genes are used as probes to determine the number of copies of the LA gene in the genome. For example, some cancers exhibit chromosomal deletions or insertions, resulting in an alteration in the copy number of a gene.

In another preferred embodiment LA genes are used as probes to determine the chromosomal location of the LA genes. Information such as chromosomal location finds use in providing a diagnosis or prognosis in particular when chromosomal abnormalities such as translocations, and the like are identified in LA gene loci.

Thus, in one embodiment, methods of modulating LA in cells or organisms are provided. In one embodiment, the methods comprise administering to a cell an anti-LA antibody that reduces or eliminates the biological activity of an endogenous LA protein. Alternatively, the methods comprise administering to a cell or organism a recombinant nucleic acid encoding a LA protein. As will be appreciated by those in the art, this may be accomplished in any number of ways. In a preferred embodiment, for example when the LA sequence is down-regulated in lymphoma, the activity of the LA gene is increased by increasing the amount of LA in the cell, for example by overexpressing the endogenous LA or by administering a gene encoding the LA sequence, using known gene-therapy techniques, for example. In a preferred embodiment, the gene therapy techniques include the incorporation of the exogenous gene using enhanced homologous recombination (EHR), for example as described in PCT/US93/03868, hereby incorporated by reference in its entirety. Alternatively, for

example when the LA sequence is up-regulated in lymphoma, the activity of the endogenous LA gene is decreased, for example by the administration of a LA antisense nucleic acid.

In one embodiment, the LA proteins of the present invention may be used to generate polyclonal and monoclonal antibodies to LA proteins, which are useful as described herein. Similarly, the LA proteins can be coupled, using standard technology, to affinity chromatography columns. These columns may then be used to purify LA antibodies. In a preferred embodiment, the antibodies are generated to epitopes unique to a LA protein; that is, the antibodies show little or no cross-reactivity to other proteins. These antibodies find use in a number of applications. For example, the LA antibodies may be coupled to standard affinity chromatography columns and used to purify LA proteins. The antibodies may also be used as blocking polypeptides, as outlined above, since they will specifically bind to the LA protein.

In one embodiment, a therapeutically effective dose of a LA or modulator thereof is administered to a patient. By "therapeutically effective dose" herein is meant a dose that produces the effects for which it is administered. The exact dose will depend on the purpose of the treatment, and will be ascertainable by one skilled in the art using known techniques. As is known in the art, adjustments for LA degradation, systemic versus localized delivery, and rate of new protease synthesis, as well as the age, body weight, general health, sex, diet, time of administration, drug interaction and the severity of the condition may be necessary, and will be ascertainable with routine experimentation by those skilled in the art.

A "patient" for the purposes of the present invention includes both humans and other animals, particularly mammals, and organisms. Thus the methods are applicable to both human therapy and veterinary applications. In the preferred embodiment the patient is a mammal, and in the most preferred embodiment the patient is human.

The administration of the LA proteins and modulators of the present invention can be done in a variety of ways as discussed above, including, but not limited to, orally, subcutaneously, intravenously, intranasally, transdermally, intraperitoneally, intramuscularly, intrapulmonary, vaginally, rectally, or intraocularly. In some instances, for example, in the treatment of wounds and inflammation, the LA proteins and modulators may be directly applied as a solution or spray.

The pharmaceutical compositions of the present invention comprise a LA protein in a form suitable for administration to a patient. In the preferred embodiment, the pharmaceutical compositions are in a water soluble form, such as being present as pharmaceutically acceptable salts, which is meant to include both acid and base addition salts. "Pharmaceutically acceptable acid addition salt" refers to those salts that retain the biological effectiveness of the free bases and that are not biologically or otherwise undesirable, formed with inorganic acids such as hydrochloric acid, hydrobromic acid, sulfuric acid, nitric acid, phosphoric acid and the like, and organic acids such as acetic acid, propionic acid, glycolic acid, pyruvic acid, oxalic acid, maleic acid, malonic acid, succinic acid, fumaric acid, tartaric acid, citric acid, benzoic acid, cinnamic acid, mandelic acid, methanesulfonic acid, ethanesulfonic acid, p-toluenesulfonic acid, salicylic acid and the like. "Pharmaceutically acceptable base addition salts" include those derived from inorganic bases such as sodium, potassium, lithium, ammonium, calcium, magnesium, iron, zinc, copper, manganese, aluminum salts and the like. Particularly preferred are the ammonium, potassium, sodium, calcium, and magnesium salts. Salts derived from pharmaceutically acceptable organic non-toxic bases include salts of primary, secondary, and tertiary amines, substituted amines including naturally occurring substituted amines, cyclic amines and basic ion exchange resins, such as isopropylamine, trimethylamine, diethylamine, triethylamine, tripropylamine, and ethanolamine.

The pharmaceutical compositions may also include one or more of the following: carrier proteins such as serum albumin; buffers; fillers such as microcrystalline cellulose, lactose, corn and other starches; binding agents; sweeteners and other flavoring agents; coloring agents; and polyethylene glycol. Additives are well known in the art, and are used in a variety of formulations.

In a preferred embodiment, LA proteins and modulators are administered as therapeutic agents, and can be formulated as outlined above. Similarly, LA genes (including both the full-length sequence, partial sequences, or regulatory sequences of the LA coding regions) can be administered in gene therapy applications, as is known in the art. These LA genes can include antisense applications, either as gene therapy (i.e. for incorporation into the genome) or as antisense compositions, as will be appreciated by those in the art.

In a preferred embodiment, LA genes are administered as DNA vaccines, either single genes or combinations of LA genes. Naked DNA vaccines are generally known in the art. Brower, Nature Biotechnology, 16:1304-1305 (1998).

In one embodiment, LA genes of the present invention are used as DNA vaccines. Methods for the use of genes as DNA vaccines are well known to one of ordinary skill in the art, and include placing a LA gene or portion of a LA gene under the control of a promoter for expression in a LA patient. The LA gene used for DNA vaccines can encode full-length LA proteins, but more preferably encodes portions of the LA proteins including peptides derived from the LA protein. In a preferred embodiment a patient is immunized with a DNA vaccine comprising a plurality of nucleotide sequences derived from a LA gene. Similarly, it is possible to immunize a patient with a plurality of LA genes or portions thereof as defined herein. Without being bound by theory, expression of the polypeptide encoded by the DNA vaccine, cytotoxic T-cells, helper T-cells and antibodies are induced which recognize and destroy or eliminate cells expressing LA proteins.

In a preferred embodiment, the DNA vaccines include a gene encoding an adjuvant molecule with the DNA vaccine. Such adjuvant molecules include cytokines that increase the immunogenic response to the LA polypeptide encoded by the DNA vaccine. Additional or alternative adjuvants are known to those of ordinary skill in the art and find use in the invention.

In another preferred embodiment LA genes find use in generating animal models of Lymphoma. As is appreciated by one of ordinary skill in the art, when the LA gene identified is repressed or diminished in LA tissue, gene therapy technology wherein antisense RNA directed to the LA gene will also diminish or repress expression of the gene. An animal generated as such serves as an animal model of LA that finds use in screening bioactive drug candidates. Similarly, gene knockout technology, for example as a result of homologous recombination with an appropriate gene targeting vector, will result in the absence of the LA protein. When desired, tissue-specific expression or knockout of the LA protein may be necessary.

It is also possible that the LA protein is overexpressed in lymphoma. As such, transgenic animals can be generated that overexpress the LA protein. Depending on the desired expression level, promoters of various strengths can be employed to express the transgene. Also, the number of copies of the integrated transgene can be determined and compared for a determination of the expression level of the transgene. Animals generated by such methods find use as animal models of LA and are additionally useful in screening for bioactive molecules to treat lymphoma.

LA nucleic acid sequences of the invention are depicted in Table 1. All of the nucleic acid sequences shown are from mouse.

TABLE 1

TAG #	SEQ. ID NO.	SEQUENCE
S00001	1	AGCAAGCAGGGAGCCAGCTGCGGGCCAAGGAGGGAGGGNGACTTTCGGTAACCGCACA GCANCCGGCGGGACAGCAGCGGAGTGTAGGGCAGCGC
S00002	2	CCGGGNTTTAAAAAGCACCGC

TAG #	SEQ. ID NO.	SEQUENCE
5	S00003	CTGGAGAGCATNTTCAGGGTGNACAGGGCNGCCNGGGCNGGTGGACAAAGGTCAG GANNCANTCGATNTAGCCCANATGGTCCTTCAGTCACAGAGCCGGAACAGGCAATTCT CTANCCATAAACAGCCACTCAGGCAGCCCCAAACCACACGCATGCACATGTGAAGACT CTGATGAAGTACAGCTGCT
	S00004	GGAGCTGTGGTCGAGGCTGGTCCAGCATATCCCTGGAGACTAGAACTGTGCAGTGGGA AATGCGGTAGACTCTGAGTTCTGGAACCTGTTGAATCTCTGTTGAATCTCCGTTTC CTCATCTGTAAGAGGTTAGTAAGTTGTCTAAGGAAAGGT
	S00005	AGATAAGAGCTAGGAGACACCCACAGCTGGAAAATCACCAAGTTCTAAGACCAC
	S00006	AAAACATGGGATTAACCTTATAACCCAGGATCAAACCTGGCTCGGTCCGCTCTTGCGG TCATCTTAGACTTGTGTTTCCTCCCTAGGAACCTCCTCAGCATGCTTTCTAA AAGCACTCCAGTGTATCTGCAC
	S00007	AGTGGAAAGATGGGATTCTTAGCCAAGACCTGATCAGGCTACACTGCCCTCGTTCA CCTCATCCATTGCATGGAGGTGACTTGGGGTGGCTCCTGACANTATCCCTCCTGCA ATTCACTCCCCATAGAGAAACTGCCAATTGCCAGTTAAGACACCTCTGTTCCCTG CGGGGCATAAGTCCATGCGCTGAGCCGGTCACGTGACNGACCTCCAACGCCTCATCC TGCTGTCTCAGTCT
	S00008	CCCTGACAGTATGTNGTGTGGTTGGTAAANACNTANCCTGTGGGTGTGGATTGGC TTAGAANGTGCATCTGGTATGTGCCAACAGGCTTCTAACTGTNCCTACNCGTCTATG TAC
	S00009	CACCCCTGTATCGGTCTCGCCACCACCACTACCAGCATCCCCAAAGAAGAAAA TCTCCTCCGAAATGCCCGAATGAGTGTGCTGGCTCTGAAGCCGTAGAATT CGTAATGGAATGTGAACTGCTCGTCCGGATCTGGCTCACGTTCTATCTTAAACAG TAAGGAACGAGGGAGGGCAAATCTGCTGAGCAAGGAAAAATAACTTCCCTCTTTT ATAACCCATCAGGATGCCGCGACGAGGGCAGCTAGCAAC
	S00010	TNATGGTGGCCCCNGACNAGGTCCCTACCTGCTTGACCTACACTTGTCCCTGGCCG CTCTGTCAACCTGGCCCGCTTGTGAGGAGCCTTCAGGTGAGGCCAGGCTGGACTGG GCTTGGGTCCCCATGGACCATGGAGATCATGAGCAGGCTGGGTGCAGTGGCTGACC ACAGGAGATGTCTGCTGGGTCTGACCGTACGCCCTGGGTGCTGGCNNTACCCCTGGG CTATTGTNTGCCAGAGTGGGGGCTGGTGCATATAACTCTAGCCTGTATCTGTT
	S00011	GGAGCAGTCATCATTGGAAAATGAGAGAAGATGTCTTAAAANGAGCCCAATCTGA GGTGTGGTGCACCTCTCTGCTGGCACACCTTACCCGAACCTCCGCGTGTGCTG CTGTCCTGGACCTTACTTGTACCTCTACTTCTGTTCTGTGAGGACTGCCACCCAGC TCAGGCCACCACCTCTGCCCCACTGTGATGACACAGAACTGCGC
	S00012	CTCGTTCAAGGGTTGCTTANAGGATTCTAAAAACAGACAATTNAGCANTCCATGTT TACCAANGGCAGTTGAAATCCAGTTCTAAATCACTGTCAACTCTCCNACACTTTC TATTGT
	S00013	CTCCGTNGGGAGCCANCNTGGACGGNGTGTGGGACCGGTNTCCAGTCNTCTCCGCA AANCGGTCTCCNAGGTGGTTAACCGGNNTTGGTGGNGGTGGGTTCTACAGTT GATGTCANCTCANCTAGTGTGACATCACCCCAAACCAAGTGTGATTCTCCCCAACAT CCCAATCACATCCCAGCGATTGGCAGCGCAGGGAGACATTGACTACCTGGGGATGA CTCTGAGGGTTAGAATTCTCAGTTTACTTAAATTGTTGCTGCCATGTCGATTTC AGGGCAGCNAGGGGNATTAGATGCCCTGTCCTTNGA

TAG #	SEQ. ID NO.	SEQUENCE
S00014	14	ACTTCACCGANATGTAGCAAGAATTCAAGACGGATGGG
S00015	15	ATCTCATCTCATCTCATCTCATCTTCTTCCATACTTATGTTGCCTATTCAAGG AATATTTGGCTATTGTACCTGTGGATATTCAATTACAAGGAGGCAGTGGCTCAAATG AAGCCAAAGAGCCTGGCTCTGAAGGACTGATGCCAGGTGGCCAGACATAGGTATTCA AAANAAGATTTGAGGCTCTGTTACCTCTCGCTGATGGTGCCTACTGCTGAAGTAGT ACTTCTTACCTGGCAGCATTGTCTCAGTGACAGCTGTCTTGTCACGGGGCCTC TGTGCCCAGTGTCTCACAA
S00016	16	TCTTGGANGCTCNAAAGCTTGCAGGGNGTTGGTGTATCCATGGCAGGGACTTGAGTTG ATTATTTTACCCGCAAACAGGGTANTGCTGACCTCGAACCTCAATCCTTTCCCC AAGTGTCTGGATTACAAATGTTGTCTACACACCCAAACAAATTAAATGATNCAAGA ATTNTCCCCGTGGCC
S00017	17	ACCCAACACTGCCATGCCTCCCCAAGCCAGATTAAACTCTTCTCGATTGCCTT TATACTTCTCTACTCTCGATAATCCAGTCTCAAGGCCCTAGAGAAGGAATGACTG TGCGTCCCTTTAATTTTACCCCTAGAACCTCCCTGATTTTTAACTCAGTGACCAC
S00018	18	AAAGTGCCAACTCTGCAGNTGNTCTTCACTCCACCACACTNGGNNTNCCTGACTG GCTACAGAGATGGAGTCTCAGNCCAGCTCCGCCAG
S00019	19	TTAGGACTGAAGGAGCTGAAGGGGTTGCAACCCCATAGGAAGNATAACNATATCAAC CAACCAAG
S00020	20	GAGCCACACTGGNAAGTCTGACAAGAGTCAGTGCTGTCCATGCTGACTCCACCCCTG
S00021	21	CTATAATGATATACCAAGATAAAGGTAGAAAGGGTGGTAGTCTTTATGGAGTATGT TTTGGGTTAAAAGTTTATTTGATATTAGAAGAGCTCAATTCAAAACTGACTT TTAAGGCTCAAACATAACAGAGATAGATAACCAGTATCCTGTAAATGATCAAATAAT TTAATCTGTTAGAAATATATAAGAAGCCATGCTAAGAACTGATGCAGTTAATTCAA GATTAAGCTTATTTAGTCTCTGTTGATATTTCAGGTATAGTTAGACCAGATA ACTAAAAACAGGTAGGTACTAGCCCTCAAACCAAGTCAGAGATCTCCTGAATGTGGCAT TTAG
S00022	22	CTACTTGGATCTGATGATGNTGCCAGGATAACAAGAAGAGACACAGTCAGCCAGTCCT AAGACAGACAGACTTCCTAGGAAGCCAGTGACTCTCAGCATGAAAGGCACCAAGNACT GGCAGCCAGGACTCAGGNCCTCTGGCATTCTGGCTACCTCCCTGTCCCC
S00023	23	TNAAAAGATTGGGACACCCCTCCGGGGCCACGGCCCTCCGGGGAAACCA GGCCCGCGTCCTCTAGCTCTCAGGCCAGGGCAGAAGTCCATAGTAGCCCCGATCAAAT AATTATCCCGAGCTTGCTCCCTGGAGGGAGGTTAAACCAAGGGCCCTGTGCACACTAC CCCGATGGGCACAGGCAGG
S00024	24	CNTCTGACCAGCTCTAAATGGCTCTNATTACNTTCAATGGAGCATAGAGTCAAATT TGACAAGCACATAAACTTAATAGCTGATCTGCAGGCATAATTACCAACAGACTGATT GTAAC TGCCAGCGAATAAGCCCACGGAGACGGTTATCCAAAGTCTCCAGTTCAAAGAC CGAAGTTGTGAGGATGAAGCCACTACAGCCACGTTGGAGCTAAGCGTCTGCTGCATT GAGGCTCTAGACACAAATGCAGGGAACTGAGCCATCTCAAAGCATCACTC
S00025	25	GTTCAATTAGCCCTGAAAAACTACACTTCCTCGTGG
S00026	26	TCTTACCAAAACACAGCTAGGGNCTCCAGTGGANAACATTCCAGCTGTGTTTTGCCTG ATTGCATCAAAGCTAGGGNCTCCAGTGGANAACATTCCAGCTGTGTTTTGCCTG ATGACACACACACACATAGATAT

TAG #	SEQ. ID NO.	SEQUENCE	
S00027	27	AAAGGTGCTTCTTAGAGGTGCTAATTGGGAAGAGCCAAGGTGAAGGCTGCAGGACACA AATGTATCTCTGTGAAATCTGCTATGGAAACATCGTCTGGGACCTGTTGGTGGAAATC CTATTGGCCTTGAGCAAAAAGGCGAAA	
S00028	28	TTAAAAGAACCCCTGGCTTCCCAGTTCTGCCTCAGGCAAAGGAGCCTGCTTACATTCC AAGCAGGACTTGTGCCCTCCAGATAGGGAACCCCAGGAAGGCCACCGCCCGTCCCAGAC CAATTCTTCCCTCCCTTCAGCTCGGTAGGTCTTGCACTAGGATCCCCGCCCCAGA CCGCCTGTGAGCAGAGCAAAGCGGTCCCAGCAGCTCTCAGATACTGCTGTGGTCTG TGTCTGCGAGGAAGGCAGCACAGAAACTTCAGTCCCCGGGTATTTGTCAGTGTGGC TCTTTATGTTACCGCATCCCACAGGGAGACACGGTTATGCCATTTTATTATCTCTC TCCCCTGCTGGGAGCTTCTTC	
S00029	29	ACAGAAAAGAAGTCTGGTCACAACAGGCTACAGCAAACGAGCCAGGTACCCAGGGACG ACTCNCCANTTCCNGCCAGAGATCTGATCTACGTACACCTGCGTCATGCTGAGACCC CNAGCCTCACTAAAAGGTCCCTGCCTAGTTCTGTTACNAATCTGCCTTATTCTGTT TTTGGTCCCAGTAAAGATAGAGTNAATACCGTATT	
S00030	30	TGTGAGCAGAGGGTTAAAGACATGAAATCTGGGCTGCAGAGACAGCTCCATAGTTNG CAACACCTGCTGCTCTCAAGAGGACCCAGAGTTGGCTCCAGCACCCACATCAGGT NGNNNANNNGCACCTGAAACCACAGCTCTAGGGGCTCAACCTCTGGGCTCTGCAG CGCCAGCATATGCACTTGCACGCG	
S00031	31	GGTTGCGGTCACATTGGCGTGTCCCCAGCCGGGGACGGGGCCCCGGGAGGCCCC GCATCGCTGCANT	
S00032	32	CTTGCAAGAGTNATTGTTGCTCCTTCTACCANTTCTAAAGATNAGACGCTGGTTG TCAGCCTCTGGCAAGC	
S00033	33	GATNNCCCANTATTCACTCTGATAGTGAATATACCCAAACATGACACCACCCCTCCGGG ACAAAGGAAGCACATGCTGGCTTGCTGGACCCCTTAAGTCTGCCAGCTTAGGTAN GGACTTCCTGTCCTCATNCACTGGGGAAAAGAAGTGTGGAGAAACGTGTACCANTA GGTGTGCCCGACAACGGTCTCGATCAACCAAACAAACATACAGATCNCTC	
S00034	34	ATTCCACAGGTAGAAATGCCACATCTTACCTCATGTGTTGCTATACTAAAATATTCA TGCATTGAAAATACTGTATGAAGCCGGCAGTGGTGGCGATGCCCTTAATCCCAGCA CTCGGGAGGCAGAGGCAGGCAGATTCTGTGAGTTG	
S00035	35	CTATAATGATATACCAAGATAAAGGTCAAGAAAGGGTGGTAGTCTTTATGGAGTATGT TTTGGGTTAAAAAGTTTATTTGATATTAGAAGAGCTTCAATTCAAAACTGACTT TTAAGGCTCAAACATAACAGAGATAGATAACCAAGTATCCTGTAAATGATCAAATAAT TTAATCTGTTCAAGAAATATATAAGAAGCCATGCTAAGAACTGATGCAGTTAATTCAA GATTAAGCTTATTAGTCTGTGTTATTTCAAGGTATAGTTAGAGCAGATA ACTAAAAACAGGTAGGTACTAGCCCTCAAACCAAGTCAGAGATCTCCTGAATGTGGCAT TTAG	
10	S00036	36	GCTGAAAATGCTAGGCTTGTNGAGCTATGAGCCCCGGGAATCCTCTGTCTACTT CTCCAGCNGAAGGATTACAAATCTACTCCACCTGAACATGGGTGCTGNAGGNGAAC CTTAANCTCACGGAAGNTCANCAGCATTNACAAACCTGTCATGCCCTGNTTGTGTT AAAGATTNATTATTGATAGGCATGATTGTTGCCTGCATGAATTCT

TAG #	SEQ. ID NO.	SEQUENCE
S00037	37	CTTTAACCGTCCTCTCCTAAAAAATATAAGAAATGAGTAAATGGGTGACTGGAGGAAC AAGAGAAAATAATAGTGTGTAANAGGGTGAGTCCTCGCTGTTGGTCAGCACAAACGCACC TGCAGAGGCTTCTTCTCTTTATACGTTAATAATGCTGCTCCATCTCCCAGGG ACGTTGAGGCTCAGCCTACCAATGTTCTCCTCTGTTCTCCCCTAGCCTACCC ATCACCACTCACCCCTGCGGCAGCCACACAGGCCCTCAGCTCTGTTCTGA TTGAATCGAT
S00038	38	GTCTCTCCTGCTTGTGAAGTAGCTGTTGTGTCNCCTCCCCCANCCCACCCCTCAAGC TCACACAGATCCTCCGAACATATGAAGCAGAGGAGGGGCTTAGGCTGCGGAACCTCCC
S00039	39	GTCTGCTCTCCTTCCCAGCTAATCTAAATATAAAAGAGGACTGCAATGCCATGGCG TTCTGTGCTAAAATGAGGAGCTTCAAGAAGACTGAGGTGAAGCAGGTGGTCCCTGAGC CTGGAGTGGAGGTGACTTCTATCTGTTGGACAGGG
S00040	40	AAATGACAACGGGGAAAGATGAA
S00041	41	GGGTACGTGGCGAGGGCTGCCACTGGTGAGGTCTCTGGACCTATCGATTCCCGG CTGATGCT
S00042	42	CCATAAGCACACATATGTAAGGTTGCACACCTCATAAGCTTCACTTTGTGAACGT GTACAGCGTTAGTATGTGAAAAAATATCATGTCGGAAGAGCAGTTCTATTGTGCT ACCCAAAAACGGTTGTATTTGAGAGGGGAGAATCACGCTGTTAGGTTTATTAT ATCCAAGTGTCCCTAGCCTCTGCAAAAAGGCCAAAGCTTGTGTGCGTGTGT GTTTAATGCAGAACACGAAGGACTCAGACACTTCGGACTCTACAGAACAGAGCA TACATCGCGGGCTGTGT
S00043	43	CCCNCTNAAAAANAAGAACAAAAGCTTCTCGCTCCTACATGGCAAAACACAAACCA CTA
S00044	44	ATAAAAAACCAAGGCATGCAAAGGTGAAAGAACCAAGTCATCACAGACGACGGCC
S00045	45	CCAGGCTGGAGGGCTGGGGGACGGTGCCTGAAAGGCACCTCG
S00046	46	CCCCCTGCCCTCCGCCACCCACCTCTCCAAACG
S00047	47	ATATTATCACTACAGAACATGAGGATGTCGTTGATTGGCGAACCACTAGACCACAC TCACTGGATGAGGAGCTCAGGAAAGCTGGCCCCATTCTCACTGGCAGCAGCACAGTA GAGCTGGCCCTAGTGGCAGGGGTGTAGGTGAGCCAGCCCTGAGGGCATGAGTGTGGGA GAACGTCCCTGCCACAGGTATGCTGTAGGCTGGTAGCATGGGCACAGAGATGATTCC CCCTCCACCGCTCCTGTCACTCTGTCAGTGGGAAGGCTGCCTGCTGGTCCCTGAGC TTGGGAGTGTATCCATGATGCTGGAGTGCTATCTGTGATGCACACGAGCTTCACCA GGTAGGAGAAC
S00048	48	TTATCCCCGCGAGACAGTCGTGATGCTCNAAGTCAGCCTATCGATGTGTTACCGTG TCTTTGGTGGGGCCTGGCAGCAGGGTGGAGCAGCCCGCGCTCTGCGGCTGGACT GAGCGGGCTGTAAATTAACAAGCTGGACGACCAGTGGCACATCCAGGCTGGCTACAA GGGGTCTTCTCGGGAGGGACCACAGGGCTTTCCAACTCGGCCGATGGAGTGC GAGGCACACTGATGCGAGGCTCCACTGCTCGGGCGAGGCCATCTCTCAGTGA TTGGGAGGACTCGCCACGTGCGGGAAACTTAAGCAGAGGCCCTCATTCTACGATGAG TGGTGCCACCTGAGGGTGGCTTGGCATCAGGCC

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10

TAG #	SEQ. ID NO.	SEQUENCE
S00049	49	GGTTCTTGAAAGAGCAGTCAGTGCCTCCATTGCTGAGATATCTTCCAGCCCCCTAT TTTAAANATTNAGACAGGCTTCAAGGGCTAGCTGAAACTCACTATGCAATAGAG AAGGACTTGAACCTCTGATCCNCCTGCCCTCACCTCCCAAGTGCTGGGATTACAGCCC CCACCCCCACCCCCAATGCCAGTTGTATACTGTAGGCAGTGGAACCCAGGGCTCCAG CATGCTGATGCTGGTATGCATGGGACTTGGACCACATGCC
S00050	50	ACAGAAAGGAAACCGATTGTTCACTTGGAAATTCTCGAATCTAAT CCAGCGTTAACTCACCCTGAGAAGAGCGCTTGTCTCATAGGAGGCTGNGTTAA
S00051	51	AAATGTTTTGGTTTTAAATCGGGCAGGGTGCTGCCACCTTAAATCCCAGAAA GAGGAAAGCAGAGGCGCGTGGCTCTCCAAGCAAGCCAGGCTAGTTCCATCCATCTG CGGGTTATCCAACCAGAGAGAATTCTCTCATTTGGTTCCGACATGCTTAGGCAT AACCTGGGAACGAGGGTAGGAGGGAGCTCCAGGCTTAAGGACAAAGGAACCGCAGGT GCAGGAAGCTCAAGGAA
S00052	52	GTTTCAATTAGCCCTGTAAAAAACTACACTTCCTCGTGGCCG
S00053	53	TTCATAATCTGAGGCCAGCGTACAGCTATAGAGTGAGATCCTATCT
S00054	54	AAAGTTCTCTGAGACGTGNNGACTCNGGGCGTGGCAAGTGCNTGTTGAGTGGATC TGTCAATCCGTTGTGTGATAAAACTGTCAACAATGAAGGGATATTATTTAGCTTATAG AAAGTCCTGAGCCANGAACTGAAGAGGGAGGCACGCACTCATGGCTAGGANGCAGCTG GCTCTGGCTGGCCTTGTCCATCCTACTGGGACT
S00055	55	CCACTCCCCCCTTGGCCCTGGCGTCCCCTGTACCGGGGACACAAAGTCTCGTG TCCAATGGGCCTCTTTCCAGTGATGGCCACTAGGCCATTTGATAACATATGCA GCTAGAGTCAGAGCTCAGGGTACTGGTAGTTCATATGTTGTCACCTATAGGG TTGAAGATCCCTTANCTCCTGGTACTTCTCTAGCTCCTCCATTGGAGCCCTGT GATCCATCCATTAGCTGACTGTGAGCATCCACTCTGTGTTGCT
S00056	56	GACGGTGTGCACTGAGAATAAAGGTCTCAGCAGTCAGTGCAGAAAATCAAGCAAAG CCCCCTTAGGAGTTATTGATGTTGCCGTTCTGTGCAAATAGGGAGGGGCTTAAG GCTTACCGGAAGACCCCCCACCTAGCTCAGGTCTGTACTCTGTCTGGTAAAG GCAAAAGGAGATTGGGGTAGTTGATGCCATTAGGGGGTCTCGCAGACTAGA AAACCTGAAATGCACTTAAC
S00057	57	AGGAATCCAGAGTTGTACACAGCGAGGTCTGAAC
S00058	58	AGAAGAGTTGGTAAACTCATAGAAGCCCTGAAGTATTGTTAGGTTGGCTGCCAGT TTAATCGTAATTGCTGCTTTCTACAGGTTTGCTGGTGTGAAATGACTGAGTACAA ACTGGTGGTGGTGGAGCAGGTGGTGGAAAAGCGCTTGACGATCCAGCTAATC CAGAACCACTTGTGGATGAATATGATCCCACCATAGAGGTG
S00059	59	CCCCCCAAAAAAATANTGTTGGAGCACCAGTTGATAAATATTGCTCAAGAAATT TGCCCCGAGGACTTGGAGCTGACAGAAGGTCAAAGCGAAGTGTGTGATTATGTTCTC CTGACAAGATACTGGCTGTTACAGACACAAAGTTGAGNCTCCACGGTCCACAGA CA
S00060	60	CTATGTTGATCTGGATATTAATTACAATATNCAAAACAAAGCTGGTATATAGCCT AGTGGTAATGTACTGACTTAGCATGCCGAAGGCAGGCTGGTCTTTATGAACTTA CAGCCTGTCGGTTTATCAGGATCAGCACATACAGCTGGTATCTGTGTCTGTGGAACT GGTAGGTTGAGACTCTTCCCCATGGGCC

TAG #	SEQ. ID NO.	SEQUENCE
S00061	61	AAAAAAGTTCTAATTATCATGTGAGGAAGANAGTAAGTTATGAGCAGCCTCTGGAAAG CATNGCAGCGCCTCGCTCTGCTCCCTCTCTCTGTCTGGGTGAG
S00062	62	TTCTCTCCNCTAGACTTCTGGGGACTGGGAGACTGCAGTATGGGTCGTGCAGGATTGG AGTGATATACTTAGCAAGCCTCCAGCGTCTGGGTCTGCAGTGCACCTGTGCATTCC TACAGTGNTTGCCAGAACAAATTGAAAGTGGTTGAGGCTTGCCCTGCCCTCTCCAG AGCAAGGTTATAGAAATTTCAGACAATATGGCAGACACCTGCCACGTGGATAAATTAC AAGCCGGAAGATTGCAATGCTGCACTTGGGTTTTGTTTGTAACTGTGTGG GATAGTTCTGCACATGGTCAGAGGCAAATAAGTCATTCTGTTGGTTTTGTTGAA GGCAAGGTTCTCTGTAGTTCTGCTGTGACTCAAAACAGAACATCCACTCACCT CTGCCCTGAGTGGTGGGATTAAAANTGAAGAACCCCTCATAAGGC
S00063	63	CTGTTNANATTAGAAGCTGAACCTCCAGCAACCACCTAAAAGCCAGGGTGAAAGAT GCATGACCATAATGGCAGCAATGGGATGCAAGACACCTGAGAATCCCTGCCAATCAG GATAGCAGAACATCCATAAGCCTAGACTCAATGAGAGGCCAGAAAATAAGGCACAG AACAAAGAGAGGAAGACACCCAGTGTCAACCTGGATCTCAGCAGGTT
S00064	64	TTTGANTCAGGATGTGCATAGCTCAAATTGGCCTTAAATTATGATCTCCCTCCTCA GCCTGCCAAGTAACTAAGATTATAGCCCTAAAACACCAGGCCCTAGGTATAAGNATT GTTTTCTTCTTTTTTCNTTTGGGTTGTTTGTGTTGGGANACANTGTT CTCTTGTANCCNGCNNTNTNTT
S00065	65	ACCAAGAAGAGTAAGAGTCATGAGGGCAATTAGAACACTTGTGTTCAGCACTGGTC GCCGAGGCTTAAACGACTGCAGTCAGCTAACTAGGGATGTCGTAGTTGTCGCATCGG ACGGCACTCCNNNNNNNCTAGTTCATCATCATTGAGCCACACCCGCCACGC GCGGCGCCCCGCGATGCAAGACCTGACTTACCAAGGCTCCCTAGATCTGTCAGC CAAGACGGAGCTGAAGAGGCTGGGCCGGCTCAGCATCGCTCCAGAACCGTCAC C
S00066	66	TGTCCAGGGNATTCACTCAAAGCGCTCAGTNCAAGCTNGCCAANAATNCTGNATAAG CGNTCANTTCAAGNTNTCCAAAATTNCNGG
S00067	67	GGACCTCAGCTTCAGAGTCTGTTCTCCCATTCTGTGGGTCTGTGAACTCAAGTN AGCTCTCAACAAGAGCAACAAGAGCCTTACCCGAGAGCCATCTGACACCCCATCA GTCATTTTTNTTTATTATTGGAGAAACTAACCTGCTGGTCTGGGTGCCCT AGCCTCTGGAAAAACTCCTACAAAACCTTCAAAACAATGCAATAAGGAGTGGAGGG ATTCAAAAAGTCTCGGGCGCTGGTTGGCTGGAGGCNATGCACTGCGGCTGGTCAG TGGGTGGC
S00068	68	GCANTTAGGAAGGCAAAGGCNTGTNATCNTAACATAATGAAGGTAAGTTAGTTATA GAAGGAAGTAGTCATGTTGAAAGAGACGGNTANTTGAGCGGTAGATAAAGTAAGAA GAGAAAGATTG
S00069	69	TGTAGTTAATAACCTGGTAATCCCTGCTACCCCCAGGGC
S00070	70	GAGGAGAGGCTGTCNCNTGGATGAGGTGGATCATNTGGGTCTAGACGTGTAGGT GGAGAGCACAAGTCTNATTCTNNNG
S00071	71	TCTTGTNTTGTNTNNNGTTGATGATNTTGTGAGTNNGANNNGGGCCTGGNNNTNNCG ANNTNCTGTCTTGATTNNTGGAGCGGGCGATTGAGANTTCGAGGCCNNNGAGTNN ANTTNNNNNAGGGATTATNNGGGANCTNTGATGGTGGATATNNGGGTGGT

TAG #	SEQ. ID NO.	SEQUENCE
S00072	72	TNACTGAATGGGANCTGGGCCAGAGGCAGTTGNCTNTGNAAAGTNCGGGTCTCA GCTCAGAGCCCTAATCCCACACTGGCGCNACAGTCAGCCGGTGGAGCGAGATAAAC GGCAA
S00073	73	TTCTGGAAACTGAATNAAATNTTTATTACAGTGATTNNGCNTCTCTGGATCTATT GATTGAGTTGGTGATACTGTTGGATCACGGGATTAGGCCAATGGGACGCCCGN CNGA
S00074	74	TGATGCTAGGCNGGCTTTGCCAACTGAGCCACANTCCTNAGGNTNTCTGTTNGG GTGCCTGGGCTGTCCTGCCAACCAGGGAAATCTGGANTCCNCGGGAGGCCAGCTGN GCTGGGACACGCTCCAAGTCNGAGACCACNAGCNGNGATGTNGCNCG
S00075	75	GTNNCTTACTATAGGGTTTTATTGGTAAAAACTTCCCTGACTTGACCAAACTTG AAATCTACAGCAGTTAATAGCACATCAGTGTCCCTGTTAGCATGGTCAGTGTAC CCTGGTTCTAGGCTTGGGCTTGCAGATGAATCAGCGTGTCTCTGATTCTGCACATT TCTGACGTGTACCGGC
S00076	76	AAATGTTTATTGTGTGATTNGGTTNTGGATGTATTGATTGNGTTGGTGATA NTGTTGGTNNGAANTGGGTGTGCNGNAGGGANGTT
S00077	77	CAACNATTACCGTGCNNAAAAAAATTTTTNAGNNTTATGCGGGGNCCCCAAAAA AAAGGTNTTACTGCTGTTATTNTGGANNTATTAAGTGGCTNTTGGTT TGNGNTATTGNAACTTTGGATNTGAGTATGTNAGTGTCTTGGNTAAGTTTG TGTGAATTNTNTTATATGTGTCTNACATGTGTAGNNAGTNGAATAAAATGGAGATTG TANGAGGAGACANTGCGATGANACNANTGGTAGNANAGNGTGGTGTGTTGATTGCAT NTTGGGATGGACTGATTTGAGTNAGATTNGGANTGGTAGTGGTGGTTAGATGCT GTGGAGAATTGGGATGGCCTTGTGATGAGGATTGGATTGGTTAGNAAAAN GATTGTTAGANTTTAATTGTGTTCTNTTCNCNGGTGGTGTATNATTGAAAGTGTATT TTGGGTNAAGATTTGGANTGAANTGTGGAAAAAAAT
S00078	78	ANGTTTTGTGAATTGATGGANATGNTGANTGGTGATTCCGNTTNTCTGGATT TTTGATTNGTGGTGATANTGTTGGTNAG
S00079	79	GCAAGGACATACATCGGGACGCTTCAGACTTCCCACTCATACCTCACAGCTCAGGGA CCCAACAGGATCCTCAGAAACACAAGTCTGGTACCCCTGCCTAGAATCACTACGGTGC TGTT
S00080	80	TGGTGTACCATGGTGTGACTCTAGGGGCTGTACTGTGTAACAGGGCCTTCCCTCC ACAGTGACCTGCTGTCTGTATAGTCTGTCTTTGGACATGACTGTGCTGTGG AGAGCAAGATGGCTGGGCTGCCTCTGGCCCCAGCATGTGGCAGCTGTATGGCTG GGGACAGACACTTTGCATCCCTGTTCTTCACTCCAATAGGC
S00081	81	CACTAGAGACCCGTGTCCAGGTGACTCTGCCAGGGCTACAGAACCTGGAGCAGGCC GCCTGGGAAGGTGGCTTCTCCAGATGGCATGGCTTACGTTAGCAACAGGCTT TCTTGCAATTTCGCAATTGCCAATTGTTGGTGGCACTCTCAAAACAAAACCTCTAGGG CTGGAGAGATGGCTCAGCTGTTAACGGCGCTGGTGGTTCTAGCAACAAGAATGGAGG TTCCNTTCTGGCACCCANACTG

TAG #	SEQ. ID NO.	SEQUENCE
5	S00082	ATGCTTTCAAAAAACAACAAAATATCCAAGTGTATTGGCCTCACCTCTGTTCT CTACTTTATTGGAAAGAGATGTACTGTGGCACCATTGACAGATGCCCTTCTGGTGGC AGGTTCTTGTTGCTGACTCTGGACTCAGACTCTGCCTGTTGCCATCTGTAATAG GGATGGGCCCTCCCTCTTGCACTTTCAAACACNGTTCTCCAAGGTATGTTCTGT CATCTGGCAAATGGGCACCTGGGA
	S00083	ATGGNTATTNTCGCGTAGGNNTNTATTCNACCAACCCANCTCCTATACNAATA NTCTGCTGCAAACGGNTCCNCAGGGCAAAGAGGATTGCCTTGTGAAANCNACT GTGGNCNTGGAACGTGTGGAGGTGTGGGTGTANACCGCANANACTCNCCCCGG AGGACNGGGTAGAGCGCCCCCCCCGAATTCCCTGGACAAAGCTTGACTGG
	S00084	TTNTCACNACGANTTGAGTATTNGTAGTGTAACTGTATTATCTGGNTTAAAAAATATTC GTNTCAAAATTNGTTNCTGAAGAANTGAGTCNTATTNTAANAAAATTGATATCNA AGGGGGGACAAAATATAAAATTCCNGGAAACANNTGACAAATACACAATAGACCGG GGNCCCCCGAATTCCCTGGACANACTGANTNGNACGC
	S00085	ACTATGCAGCCAGTTCAAGCTAGTTGAACTTGCTGTTGCCTGGACTTCCC AGTGGTCGGATGANAGCCACGCG
	S00086	GCNANAANAGGAAAGAACATTATTNGGTNGAGGTCTCCACCTTGTCAAGACNCANGT CACCACCTTGGTGACAAGTGCCTTACCCACTGAGCCATCTCACTGGCCCCGCTGT GCGTACTNGTGTGTCTGTGCGCACGCNTGTGCACNCACAGTTCACTTNAGCAT GCTGTATGTCAGCTATAGTCCTGAGCCCTCGCAGGCAGGACTGTNGCTGACCTTAC ATNTTCCG
	S00087	ACACAATGCCTCCCCGGAGATGGAGTGGCTGTTATCCCTAAGTGGCTCTCCAAGT ATACGTGGCAGTGAGTTGCTGAGCAATTAAATAAAATTCCAGACATCGTTTCTG CATAGACCTCATCTCGGGTTGATCACCCCTATCACTCCACACACTGAGCGGGGCTC CTAGATAACTCATTGTCGTCCTCCCCCTTCTAAATTCTGTTTCCCCAGCCTTA GANANACCTGGCCGCCGGACGTGCGTGACCGGTCCAGGGTACATGGCGTATTGT GTGGAGCGANGCAGCTGTTCCACCTGCGGTGACTGATATACGCA
	S00088	CTCTGGCAGCCATTGTGTTGTTACNGCANANCAACTGCTGCAGGCCTGCCTCCCT CTGAAGCTGCTGTGCTGCTGATAAAACTCTGCCCTTAGTGCTCACTGTTNCTCATAC TGTGTGCANCTGAGCAACAGCCGGATGACCATCCTACNGCAGCG
	S00089	GCTACAGCTCGTCAATGCACACGTTCTTATATAACTACACAGATCTGTAAACGA AGTCTGGACATCAAAGCTTATGGGAACTGCTAAGTGGCTAAGGACGC
	S00090	ATATAATAAATCTAGAACCAATGCACAGAGCAAAAGACTCATGTTCTGGTTGGTTAA TAAGCTAGATTATCGTGATATATAAAAGTGTGTATGTATACGTTGGGATTGTACAG AATGCACAGCGTAGTATTCAAGGAAAAGGAAACTGGGAAATTATGTATAAATTAAAA TCAGCTTTAATTAGCTAACACACACATACGAAGGAAAAATGTAACGTTACTTTGA TCTGATCAGGGCCGACTTTTTNAATTNCANANTNCAATCCCATTANTAAAAGG GNAAACCTNGGNTTTNCNGGAAGNAAGGGNTTAACGGTTCCCT

TAG #	SEQ. ID NO.	SEQUENCE
S00091	91	TTAGNTNNCTGGAACTTGNTATGTANATGANGCTTGNCTCNAACTCTGATATNCACT TGTGTCTGCCTCCTGACTATGTGAACCANACCANTCTNTNATTCAAANANACTGAGGT TGGACCATCCTTANTCACCTGGGTTGTTCTATTAANTGTAACACTACACTCATAAATTG AAGCAAANCAAACCGTACCCANCTGTGCTACTTGANGCACCTGANCAATTNACAANGG ATCTTTAACCTCATGAGGCCAGTCCTGCTAATCCAGGTTGGCTNATCCTGCAAT CCCCGTCTCACAAACACCTGT
S00092	92	GTCAAAATACTGAGAATTAGAGGCTATTGGATGCCAAGTCATAGAGAGGACACATATA TACCAATACTTCCAAGGCTCAGGAAACATCATGGAAGAAGGGTAGGAAGAATTAAAN AACACAGAAGGGGGGTGAGGTATGGAATGATGATTCCAGTCATGACTTGGCTATT GAGTTAACAAACAGCTGGATCACCTGCACAAGATCTCCACAAGAGTGGGCCATTAACA CTCTATCATGAAAGAGGAGGGCNTATGAGGTACCACCCCACCCCTGAAGATTATAC ACAATTAATANTTGGTGAGGTAGGGAGAGACATTTACTTTAGGGGTGCAGTCACTAGT ACAGTGCCTAC
S00093	93	CCATCTCTCCAGCCCCCTCTTTCTAATATGTAGGTCCCAGGGACCAGGCTCTAGC TCTCAGACTTTGCTATCTCGTGTGGAATTGTTACATTATAAGGACTTGAAGC CTCATGTCACCTGCACCACCCCTCTGAGTCTGACC
S00094	94	CAGCTGCGTTGCGTCATCCAGCCAGAGCTCAGAACAAACTATGAACTACAAAGTTCTT CAGCACCAAATCTCAGAGGCAGAAAACATTCTAGGCCTAGATTGATAGAGGC TAAGAGGCTTCTAATAGACCTAGGTTCCAGAGAGAGGTTGTAAGCCACAAAGACCAC AATTACATCAGGCGAATGAGTTACTTTACATATCTGTAAAATGAGCAGAGAAGAGTC TGGGGCTCCTCTGTTCCCCGTGGTTCCCTGCTGGCCCTGGTTTCTGTGAGATGTG CCTGACTCCCCGGATGCCCTCAACTGATGTTGGCTTAGGGGCTGAGCTTTAAATG TCAGATCTTCTCATTTCCGCTCTGTCAGG
S00095	95	AGNGGTACCGGGTANAGCANANACTANCNTACCCCTTGGGCGCCTGTGGTCTCCACAC AGAGTGTGTGGGTGTANGANACANGCTGATGGGACTGCCCTCGGCAGCCTCACGG GCACCTGTGAGTGGCAGTCTGAAGGGTGGTGGCCGGCANACANCCTATANAGTGTAT TCCAAAGCCTGAACCATTGTNGCTCCGGCTGATTCTGGTCTGCCCTGATAGTTTA GATGACCATCTTATTGTTCTTCACANGCAGTTATGCTAGANTGGATGA
S00096	96	AAACCTGTGAGCTCTGCTTTGTGCTCTACCCACAGGAGCACAGCCAGCCTAAACT GGAGCGC
S00097	97	ACAGCACCTATGGCTGTCCCTGACCTCCACACACATGTGACATATGTCCATGTATAC ATACATGCACACACACACACACA
S00098	98	GTCTCCTGGNCCTCCTGAGTCCCACACTTCTCAAACCTAAATCGGCCTGGGNCA ACATGCTCAGCCAGCAGTTAAGTCCCGTGCCTCCACCTGGAGNAGGTGTANNAAAT AGGNGGNAAGGCCAGGCCCTCGANCCGAAGGCATGAAGCCCCGGNACCGAGC ACACACTGTCTTCCCCGGGTGCCGCTCACCATCTGTTGTGACACGGGGCCGAGNCC TGAAAGNGCTGGCAGCCCCGGTGAGCGCGAANNANNGCCAAGCAGAACCGCAACA CGCCTACCCCTGAACGACATAGCAGCGC
S00099	99	GGTAAGGAANGGCTCTCTGGTTCCCTCCATGACAGGNTTCTGTGAGGGCCACGCG TCCTGTTACAGAATGGTTCCAAGTCACCGG

TAG #	SEQ. ID NO.	SEQUENCE
S00113	113	CANTGANGNNGCTCAAATGGTTAGCCTGGTGTATGTTGCAAAGGGCACTCATAGTT TACTCTGGCTTGGGCTTGGTCCCCAGGAGGGAAACAGACCCATCCANTGTGCC CTCCACNAGGTGGCTTGTTAAAAAATACCTGCNGCATTCCAGATCANCTGAGAAC CNCTGAAAAGACTTTTGTTCCTCCCTTCCAGGGTAGACGGCNNAGTCANC NTTN CNTCATTAAACAANACTGCCACCGCTATNGCTTGCCGAGCCCTACAAACCTGTA CAGC
S00114	114	AGNACCNGTTGCCAAGAGGAAGTCAAGCCAAGAAAGAACCGTGGCAANAATGAGCT GAACCGTCTGCGAACCTGGCTCGCGCACAATATGCANATGCCCANCTNGCCGGN CTGCACCCACTGGACACCAGAGTAAGGAANAGCTGGGCCGCGCATGCAAGTGGCCA AGGTTTCCACCGCTTCGGTGGGACGCTTCCAGGAGCGC
S00115	115	TTCCCTTCAGCTGTTTCAGGCATGCCACCCATCCANCACTCCCCCAACCCCACC CCGTGAATACACAGAGNGNGACAAACTCTGTGTGTGTGTGTGTGTGTGT TGTGNGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAG AGAGA
S00116	116	AGTGTATGTATACTACNTTGGGATTGTACAGAANGCACAGCTAGTANTCAGGAAAAG GAAACTGGAAANTAATGTATAAATTAAATCAGCTTTAANTAGCTTAACACACACA TACNAAGGCAAAATGTAACGTTCTTGTATCTGATCAGGGCCGACTTTTTTNN NTGNNNAAATTNCNATNCCNNNANTAAAAGGGAAAGNTNGNTTNTCNNGGNGNAA GGGNTTAANGNTTTNTTTNTT
S00117	117	AATCCTTCTGTACTGAGTGCCTGGGAGGCAGAGAGCAGAACGTCAGCCAGTGA ATACTCTCTCACCACTAGACCCCAGCTCCTGCCTCAGCCTCCCCAGCCTGGCTATCA GAGCTTAGCCCCACTCTATTCCCAGGC
S00118	118	AGTCAACATAACTGTACGACCAANGCAAATACACAATGCCCTCCCGCGAGATGGA GTGGCTTTATCCCTAACGTGGCTCTCAAGTATACGTGGCAGTGAGTTGCTGAGCAA TTTTAATAAAATTCCAGACATGTTTCTGCATANACCTCATCTCGGTTGATCAC CCTCTATCACTCCACACACTGAGCGGGGG
S00119	119	TTATNTCTCCATGGCTCCAACTGGGAGANGNNGAGGGACACTTANAATTGNCNN NGCAACNTTGAATTTCAGAAAAGANTGCTTCACGCCATGCAACATGGANAAGG ANATGGANGTGAAANTTCCATGGACAGAAAGTAANAACACTCANACNTCTNANTGA GGGCCTGAANTNTGCNTCCATTATA
S00120	120	TGNGCATAACACACCTTAGCGAAGGGTGCCTGAAATCCGCTCAGGGTAACCTAGGCGG AGCAGCCGTGTAGCACGTGGCTGCCACGCG
S00121	121	CCCCCAATTGCGATATTCTCGNGNTAGCGCTTGATTTCCCCACCCAGCTCC TAAACCAGANTCTGCTGCAAACCTGGCTCCACAGGGCAAANAGGATTGCGCTTTGTG AAAACCGACTGTGGCCCTGGAACACTGTGTGGAGGTGTATGGGTGTANACCGGCAGANA CTCCTCCGGAGGAGCCGGTAGAGCGCC
S00122	122	CTGNTGCCAGCTTAAAGCTCAAAGCTTCCACTCCAGTGCAAAGAGATGAGATTTG AATCAACAGAATTGTTGGACTTAAATGTCATTAAATTAAACTGATCTAGAAAA GCACAAAGGTGCACGTNTTCTGGGGCAGCAGTGTGTGTCAATATGCAAACCTGGGC TAATTAGACCACTCACTCACTGAAACAGAAACCACTAGATTCCCTGTGAATCCCTC TCTTCAGGAGGCCATGGGGCAGGGAGCACCCTACATCTGTGGGGCACTGGACCCC C

TAG #	SEQ. ID NO.	SEQUENCE
S00123	123	CTCCTATTCAAGTCACACCCCTGCTGCCCATANATCTACTTGAAAGAGGGAGTTAA CCAGCAAGCCTCAGGATAAGAGGACAGAAGTCACAAAAGCCACAGGAGGC
S00124	124	TGGTGAAGACTGCCAGGCTGGTCGGGAGGGCAAGGAAGGAATACAGGACGATCTGCN CATCGTATTGCTTCCAACCTGAAAAAGGAGCAGTGTGGCAACAGGCTGTTTTACA GGCTGGGATGCATTCGCCCCCTACCTGCCTGACAGCCCTGCGCACTGCAGGAAGG AGACGAAAGCATTGACCACCCCGAACGCCNAGGGAGAANGGGGGCTGGGAGCGGAC AAGACCGAAGACAGCACCCAGCTCAGCCTTCTAAGCCGGCAGNTCAGGAACCCC ACAGACAAGGGCCGAGCGACTCGTGNANCTGCCGTGGGAGGCTGTAG
S00125	125	ATCTNNNCNNCTNTGACCTGTTNNGCTCTACNTCTATTCCTCAAAACNAANNCTA GACCAAGGTNTCTGTTCANCNTNNACTTTAAGTGAACACAAATTAAANCNGNGAC ACTGGNAGAGGGAGTCACTGAC
S00126	126	GTATGGAGAGTGAATGCTTGGTGGCTTCCTGGGTGCACCCATGCCAGCGC
S00127	127	CTCAAACCTCCCTCTTGCCTCCTCACCCACTTGCCTTATNTCGAAAGCTCTT ACTCATTTCCCTTTCTGTCCTCGATGTCTCTGATTCTTCTCCANCTCTGTT CCTCCTTTCCCGGTGTCCTGTCAGGCT

Contigs assembled from the mouse EST database by the NCBI having homology with all or parts of the LA nucleic acid sequences of the invention are depicted in Table 2.

TABLE 2

			MOUSE
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
S000004	F1	128	CGGCCAGGGACTCCCCTCCAGGCTCCTCAGAGAGCAACAGGCGAAGAGAACTAAACTGTT TTGCCCTTCAAGATCAATAACCCCTCATATAACCCAGGGATGAAGGATGCTAACCCAA TCCTGCTGCCTTGTCAACCCCTCTCCCTGTTGGGACCCAGGAAAGGGCCTGGAGCAT CTTACCCCACAGGGACTCTTAAGATCACTGCCATCCCTCTAAGACAAACCTTCCC TAACATACACATTTAAGTGTGCCATTCCAGAGGGCTCTACAAGGTCTTTACCTT CCTAGACAACTACTAACCTCTACAGATGAGGAAACGGAGATTCAAACAGAGATTCAA ACAAGTCCAGAACTCAGAGTCTACCGCATTCCACTGCACAGTTAGTCTCCAGGGA TATGCTG
S000010	F2	129	ACTAGAGGCAGTAAAGTTATTACATTAACACTCAATGCTGGTCAGAGGCATCCACACG GCCCTGATCTGAATCCTGAAGGTGTGGAACCCAGAAGCCGCTGTGACTTGCAGGGTCAG GACTTGGGCTGCCTGCTTGCATAGCTAGACTCCTATGCATCCTTCAGAGGTACCCCA ATGCCCAGTCAAAGCAGCTTGTGCTGTGCCATATGGCACTACTCCTCACAGAGCA GCGCCTGTGGAGGATCTTCAACACAGCACATGGACATAGTCCCTGACGTCCACACCCGGG GCTACCAGGAAGCCCCAGGGCTGCCTGGCTCCTCACATCCTTCTCATCTTGCCT TCCTGGAGGGAGCACCCGGCAAAGGCCTGGCGAGCTCTGGCTGGGCTGGCGTCGG TGCTTGGGCTCTGCTGGAGGCATTGATCTAAAGATGGTTGTGCGCGTGCAGTAGTTCT TGATGCTGTCCACCAGCCTCAGGCCCTGGAGCTCTCCCTCTCAAAGCATGAGCTGAAGA GTGGGTGCAAGCCCAGCTCTGCCAGGTCCAGCTCCTGGCTCTTGTAGGGACTCAGGCG AGGGCGCTGCCGTGAGCGCACATACTGCTGCTGAGCGTTGT
S000013	F3	130	CCGCCACCAAACGCCGGTAAACCAACCTCGGAGACTGCTGTGCGGAGAGGACTGGGAAAC

MOUSE			
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
			CGGTCCCCACACACTGTCACGCTGGCTCCACGGAGGCCACCCACACCCGGGGCG GGCAAGATGCAGTGATCTCAGCCCTCCGCTCCGACTTCCGCTCAGTA T GGCCT CACAGCTGCAGGTGTTTCGCCCCCATCAGTGCAGTGCTGAGTCAGTGCAGTCAAAGA AACTGAAAATAGAGCCCTCTGGCTGGATGTTCAAGGACAGAGCAGCAACGACAAATACT ATACCCACAGCAAAACCCCTCCAGCTACACAAGGGCAAGGCCAGCTCCTCTCACCAGGTAG CAAATTCAATCTCCTGCTTACGACCAGGGCTCCTCTCCAGCTCCTGCCGTGGAGC ATATTGTGGTAACAGCTGCTGATAGCTCAGGCAGCGCCGCTACAGCAACCTTCAAAGCA GCCAGACCCCTGACTCACAGGAGCAACGTTCTTGCTTGAAGCCATATCAAAATGTGGAT TGAAGAGAAAGAGTGAGGAAGTGGAGAGCAACGGTAGCGTGCAGATCATAGAAGAACACC CCCCTCTCATGCTGCAGAACAGAACCGTGGTGGGTGCTGCCACGACCACACTGTGA CCACCAAGAGTAGCAGTCCAGTGGAGAAGGGGATTACCAGCTGGTCCAGCATGAGATCC TTGCTCTATGACCAACAGCTATGAAGTCTGGAGTTCTAGGCCGGGGACATTGGAC AGGTGGCAAAGTGCTGGAAGCGGAGCAGCAAGGAAATTGTGGCCATTAAGATCTGAAGA ACCACCCCTCTATGCCAGACAAGGACAGATTGAAGTGAGCATCCTTCCGCCATAAGCA GTGAAAATGCTGATGAGTATAACTTTGTCGTTATGAGTGTTTACGACAAGAACATC ATACCTGCCTTGTGTTGAGATGTTGGAGCAGAACCTGTACGATTTCTAAAGCAGAAC AGTTTAGCCCCTGCCACTCAAGTACATAAGACCAATCTGCAAGCAGGTGGCACAGCCC TGATGAAGCTGAAGAGCTTGGTCTGATTGCTGACCTTAAACCTGAAAACATAATGC TAGTCGATCCAGTTCGCCAACCTACCGAGTGAGGTCTTGACTTTGGTTCTGCTAGTC ATGTTCCAAGCCGTGTTCAACCTACCTGCAATCACGCTACTACAGAGCTCTGAAA TTATCCTGGATTACCATCTGTGAAGCTATTGACATGTGGTCACTGGCTGTAAATAG CTGAGCTTCTGGATGGCTCTTATCCTGGTCTCAGAATACGATCAGATTGCT ATATTCACAAACACAAGGCCGCCAGCTGAGTATCTCTCAGTGCCGGAACAAAAACAA CCAGGTTTTAAACAGAGATCTAATTGGGGTACCCACTGTGGAGGCTTAAGACACCTG AAGAACATGAATTGGAAACTGGAATAAAGTCAAAAGAAGCTGGAAAGTACATTTAACT GTTAGATGACATGGCTCAGGTAATATGCTACAGACTAGAGGGGACAGATATGTTAG CAGAGAAAGCAGATCGGAGAGAGTATATTGATCTAAAGAAAATGCTGACGATTGATG CAGATAAGAGAAATCACGCCCTGAAGACTCTAACCAACCAATTGTGACGATGAGTCACC TCCTGGACTTCTCACAGCAGCCACGTTAAGTCTGTTCCAGAACATGGAGATCTGCA AGCGGAGGGTTCACATGTATGACACAGTGGAGTCAGATCAAGAGTCCCTCACTACACATG TCGCTCCAAATACAAGCACAATCTAACCATGAGCTCAGCAACCAGCTAACACAGTGC ACAATCAGGCCAGTGTCTAGCTCAGCTACTGCAGCAGCAGCTACCCCTCTG CTAATTCAAGATGTCTCGCTGCTAAACTACCAATCGGTTGTACCCATCGCCAGC CAGTTCTGGAGTTGCCAGCAGGGTGTTCCTACACCTGGAACCACCCAGATCTGCA CTCAGACAGATCCATTCCAGCAAACATTATAGTATGCCACCTGCTTCAAGACTGGAC TACAAGCAACAAACAAAGCATTCTGGATTCCCTGTGAGGATGGATAATGCTGCCAATTG TACCCCAAGGCCCTGCTGCTCAGCCGCTGCAAGATCCAGTCAGGAGTACTCACACAGGGAA GCTGTACACCACTAATGGTAGCAACTCTCCACCCCTCAAGTAGCCACCATCAGGCCAGT ATGCGGTGCCCTTACCTGAGCTGCGCAGCAGGCCGGCGCTGGTTGAACAGACTG CTGCTGTACTGCAAGCCTGGCCTGGAGGAACCAACAAATTCTCTGCCCTCAGCCTGGC AGCAGCTGCCGGGGTAGCTCTGCACAACTCTGTCAGGCCCTGCTGCAGTGATTCCAGAGG CCATGGGGAGCAGCCAACAGCTAGCTGACTGGAGGAATGCCACTCTCATGGCAACCCAGT ACAGCACTATTATGCAGCAGCCATCTTGCTGACCAACCATGTCAGCTTGGCCACTGCTC AGCCTCTGAATGTGGTGTGCCCCATGTTGTCAGACAACAAACAGTCTAGTCCCTCC CAAAGAAGAATAAGCAGTCTGCTCCAGTTCAAAATCTCTCTGGAAGTCTGCC CTCAAGTTATTCTCTGGTGGAGTAGCCTCTCGTACACATCTCTTATAATTCCC

			MOUSE
SGRES TAG#	REF #	SEQ ID#	SEQUENCE
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S000015	F4	131	CCGGTCACATGTTCTTGTGATGACCACGTGATGGGTCCTAGAGGTGGGAGCAGC AGCTAAAGTCAGAGCATTGAGTATGACTCTAGCAGCTGGACACACAGAGAAATGTG CATCCCAGCTATAACTAAATCAAGAAAGGCCCTGGCTGTGAAATTACAGGGGCTCTTA GGATTACAGGGCTTGATATACCTGAAGAAGTGCACACTTTTCCCTGGCTCTCA GCCCTTCTCCAGGCTAACCTACATTACTAGATGGCTCTAGATATTCTCTCACTAAC TGAACCTTGGCATCAACACAGGCTTAAAGGACACACTTGGGTCTCTAGTGTCAATTGA ATGGCAGCATTGACTTGGCTTCAAGCAAGATGACACTGAAGTCTGCCCTTCA AACAGGGCTACCCCTGCCCTGCTTCCAGAAGCAAGCAGCCTTACCATCTGCTTAGGACT TCACAGTTCTAAAGTTCTTCCATCCCGTCTGCTTCTTTATTGCAACAAGTGT TTTTTATTGCTAGTATTACTGAGATACCGCAGATGCCACTGTGCAAGGGCGCCTGCGGT CCTGAGGAAGAGCTGTTCCATGCCCTAGGCAATTAGAAGCCATGGCTGGAATCT

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S000028	F6	133	CTGCCTTACAGCACTGTTCTGGCAGCTTACAGGAAACCTCTTCTGATTCCCACCT TACCACAAGACCCAGGGCTGTGGGGTGAGGTGTGCTACCGAAGTGAACGCCAGCAATGAT GTTCCAGAAAACATTTAATATCTCCCTGGTTCCACTGCTGCTAACGCTGGGACGGGG CTGGAATAGCCGCTCCGGTGGAGGAGGCTCCAGCAGGGGAGAGAGATAATTAAATGG CATTACCGTGTCTCCCTGTGGATGCGGTGACATTAAAGAGCCACACTGACAAAATACCC GGGACTGGAAGGTTCTGTGCTGCCCTCTCGCAGACACAGAACCCACAGCAGTATCTGAGA GCTGCTGGGACCGCTTGTCTGCTCACAGGCGGTCTGGGGCGGGGATCTAGATGCGAAG ACCTACCGAGCTGAAGGGAGGGAAAGAATCGGTCTGGGACGGGCGGGCTATCCGGGGT TCCCTATCTGGAGGGCACAAGTCTGCTGTGGATGTTAGCACGCTCTTTGGCTTGAGG AGAACTTGGGAAGGCCGGCTCCATGAGGGTGGCTCCCTTGTTGCCCCGGAGGTGGGG TTCCAACCCGGGAGGGTGGTAACGGCTAAGGGAGGCGGCTAAACAACCGGAAGGCCAAAT ATTGGATTGGCCG
S000031	F7	134	GTAAGATCTAAAGGTGGTTGACCCAACCTCAGAGCAACTTCAGGCCTTCAGGAACGAG GTGGCTTTCGCAAAACACGGCATGTTAACATCCTGTTCATGGGTACATGACA AAGGACAACCTGGCGATTGTGACTCAGTGGTGTGAAGGCAGCAGTCTCTACAAACACCTG CATGTCAGGAGACCAATTCCAGATGTTCCAGCTAACATTGACATTGCCGACAGACAGCT CAGGGATGGACTATTGATGCAAAGAACATCATCCACAGAGACATGAAATCCAACAAAT ATATTCTCCATGAAGGCCTACGGTAAAATTGGAGATTGGTTGGCAACAGTGAAG TCACGCTGGAGTTGGTCTCAGCAGGTTGACAGCCCACGCTCTGCTGTGGATGGC CCCAGAAGTAATCCGGATGCAAGGATGACAACCCGTTCAAGCTCCAGTCCAGTGTACTC GTACGGCATCGTGTGACTGAGCTGATGGCTGGGAGCTCCCTACGCCCACATCAACAA CCGAGACCAGATCATCTCATGGTAGGCCGTGGGTATGCATCCCTGATCTCAGCAGGCT CTACAAGAACTGCCCAAGGCAATGAAGAGGTTGGCTGACTGTGTGAAGAAAGTCAC AGAAGAGAGACCTTGTGCCCCAGATCCTGCTTCCATCAGCTGCTTCAGCACTCT GCCGAAAATCCACAGGAACGCCCTGAGCTTCCCTGCATGGCAGCTCACACTGAGGG ACATCATGCTTGCACGCTGACTACATCCCAAGGCTACCGATCTCCTAACTGATGATGTA GCCTGTCTTAGGCCACATGGACCAAAAGAAGTCAGCAGGACCAATT
S000039	F8	135	ACAAGACTTGAAGAGCGGTTCTGAAGAGGATTGACTTGGGAGAGGGTCACTTGG GAAGGTTGAGCTCTGCAAGATATGATCCTGAGGGAGACAAACACAGGGAGCAGGTAGCTG CAAGTCCCTGAAGCCTGAGAGTGGAGGTAACCACATAGCTGATCTGAAGAAGGAGATAGA GATCTACGGAACCTCTACCATGAGAACATTGAGTACAAGGAATCTGCATGGAAGA CGGAGGCAATGGTATCAAGCTCATGGAGTTCTGCCCTGGGAAGCTAAAGGAGTA TCTGCCAAAGAATAAGAACAAATCAACCTCAAACAGCAGCTAAAGATGCCATCCAGA ATTGTAAGGGGATGGACTACTTGGGTTCTGGCAATAAGTTCACCGGGACTTAGCAGCCA

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S000040	F9	136	TGGACTGGGTGCGGCCGGCTGCAAGACTCTAGTCGCGGCCACGTGGCTGGGGGGGG CTGCCGTGGCCCTAGTGTATTACGTAGCGGGGGGGCGAAGTGGCGCTCCCTGGCGGG GCTGTTCATGGCGGTTGGGGTCTCCAACAGCTCAGGTTGAAGTCCAAAAGCCTCCGA GGCGGGCTGGAGTTGAGGTTTGCTGGTGTGAATGACTGAGTACAAACTGGTGGT GGTGGAGCAGGTGGTGGGGAAAAGCGCCCTGACGATCCAGCTAATCCAGAACCACT TGTGGATGAATATGATCCACCATAGAGGATTCTACCGAACAGCAAGTGGTGTGATTG TGAGACCTGCCTGCTGGACATACTGGACACAGCTGGACAAGAGGAGTACAGTGC AGAACAGTACATGAGGACAGCGAACGGGTTCTCTGTGTATTGCCATCAATAATAG ATCATTGCAAGATATTAAACCTCTACAGGGAGCAAATTAGCGTGTGAAAGATTCTG TGTCCTCATGGTGTGGTAGGCAACAAGTGTGACTTGCCAACAAGGACAGTTGAC GCAAGCCCACGAACGGCAAGAGTTACGGAATTCCATTGAGACCTCAGCCAAGAC CCGACAGGGTGTGGAGGATGCCCTTACACTGGTAAGGGAGATACGCCAGTACCGATT GAAAAAGCTAACAGCAGTGACGATGGCACTCAAGGTTGATGGGTCGCCCTGTGCT GATGTGTAAGACACTTGAAGTTCTGTCTAGAAAAGAGGCCACTTGAAGCTGC ATGCCCTGGTTCTGACATCCCTGGAGGGAGACCTGTTCTGCTGCTCTGC AAGCTCCTGCTCCTGCTCCCCGACTCAGTTACTGAGCACAGCCATCAACCTG TCTTCAGAATAACTACCTCCTCACTCGGCTGTCTGACCAGAGAAATGGAC CGGTGTTCTGCCCTGGGTTCCCTAGAAACAGACACAGCCTCCAGCTGGCTTG TCTGAAAAGCAAGTTACATTGATGCAAGAGAACCAACTAGACATGCCATTG CAGTTCTTACTCTAACGTAACAACTGCTGGTATTTCCTGCCCAACTGTTGA ACTTGGCCTTGGTTGGGGAAAATGTCATAAATTACTTCTCCAAAATATAAT TAGTGTGCTGATTGATTGTAATGTGATCAGCTATATTCCATAAAACTGG CATCTGCTCT GTATTCTAAATGCAAACACGAATACTCTCAACTGCATGCAATTAAATCCA ACAAAGTGCCTTTCTAAAGTGCCTGTAGGCTCATTACAGTTGTAATTG GAATA GATGTGTAAGAACCACTGTATAGGAAAGTGACTCTGAGCCATCTAC TTGAGGGAAAG GTGTATGTACCTGATGGCAGATGCTTGTATGCACATGAAGAGATAG TTCCCTGTCTGG GATTCTCCCAGGAGAAAGATGGAACACTAACAGTAATTCT TAATCTTTTTTTTTTTTTGGTAGACTATCACCTATAAATATTG GAATCT CTAGCTTACTGATAATCTAATAATTATGAGCTTCCATTATAATGA ATTGGTCTACCA GGAAGCCCTCCATTATAGTATAGATACTGTA AAAATTGGCATGTTACTTATAGCT GTGATTAATGATTCTCAGACCTGCTGAGATATAGTTATTAG CAGACAGGTTATATCT TGCTGCATAGTTCTTCTGGAATATATCTATCTG ATGTTGAGAGAACGTGGCC AGTCCCTCTCAGCATCCCTCATCTCAGCCTAGAGAACGTTGAG CATCCCTAGAGGG CTTGAAACAGTTATCTCGGTTAACCATGGTCTAATGG ACC GGGT CATGGTT CAAAACT TGAACAAGCCAGTTAGCATCACAGAGAACAGTCC CATCC ATTGCTCCCTGCC TATT A

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20	S000046	F10	TTATAAGCCGCAGTGCCGGATGTGAATGGATTACAATGTATCTTCAGGGAAACCTATT ATTATCAATGTGACTCCTCGGGGAGTCATGATGGTGGGGAGGAGGATGATGATGA GACGCCCTCTAAACCTGGAAACAAGTTAGGACTTGTGAAAGAGAGAGAAAAAAATACA ACCAACAAGACCGAAGAACAAATTATAACTATCCAGTGTGATTATTTTATAACAAAC GAAAAAGTTGTCGGATTTTTTAATGATTACTTTGGGGGGAGGGAAATTGTTA CAGTTGATGATGGAAATGCAAAACCGAGCCAGGTGCATAATCTGTAATCTGGCT AACCTGGAACAGGACTGACTTCTATTTAAACTCTTTGGGGGAACACTCATGTGAG ACACTAAGTTCTGCAGAAGATTGCTCTCTTTAAAGTCTCTTCTGGAAATAT TGTGAGCATATTGTCGGCATTGAAGGTTGTGATTTGCTAAATGCATACCAACA GCGAATGGCTGCCTTAGGGACGGACAAAGAGCTGAGTGATTACTGGATTTCACTGCGAT GTTTCGCCCCCTGTAAGCAGTGGAAAATGGACCAACTCTTGGCAGTGACATT CACTGGCTCAAATGTAGAAGACAGAAGTAGCTCAGGGCTGGGAACCTGGGAGCCATCC AAGCCCGTCCAGGAACATGGAGATGGACTCCCTATGACCAACATGACTAGCAGGGATCT TGGGTACATGACAATCTCTCCACCTTGTCAATTCCAGAATACAAAGTAAACAGA AAGGGGCTCATACTCATTTGGAGAGAAAACGTTCAGGGTTGCCACCAGCAGACT CCTCGGAGGGACATGGATATGGCAATCCAGGAACCTTCCGCCCCACCAACCTGGCT CCAGTACTATCAGTATTCAAGCAATAATGCCGCCGGAGGCCTTCACTGAGTAGGCCAT GGAGGTACAGACAAAGAAAGTCCGAAAAGTTCTCCGGGTTGCCGTCTCAGTCTACGC TCCTTCAGCCAGCACTGCCACTAACACAGGGACTGCCAGGCTATCCTCCTCAAGCC AGCAGCCAGCAGTCCCTAGCTCCTTCTCATGCAAGATGGCCATCACAGCAGCGACCC TTGGAGCTCCTCCAGCGGGATGAATCAGCCCCGGCTACGGAGGGATGCTGGCAATTCTC TCATATCCCACAGTCCAGCAGCTACTGTAGCCTGCATCCACACGAACGTTGAGCTATCC ATCCCACTCCTCGGCAGACATCAACTCCAGTCTCCTCCGATGTCCACGTTCCATCGTAG TGGCACAAACCATTACAGCACCTCTTCTGCACACCCCCCTGCCAACCGAACAGACAGTAT AATGGCAAACAGAGGAACGGGGCAGCAGGGCAGCTCGCAGACTGGAGACGCTGGGAA AGCCCTAGCTTCGATCTTCTCTGACCAACAGAACACAGCTTTCTCCAATCTTC AACTCCTGTGGGCTCCCTCCCTACTCTCAGCAGGACAGCTGTTGGTCTAGAAAATGG AGGACAGGCCTCGTCATCTCCAATTATGAAGGACCCCTGCACTCACTGCAAAGCCGAAT CGAAGACCGTTGGAAAGACTGGACGATGCGATTGTTCTCCGGAACCACCGCAGTGGG

			MOUSE
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			CCCCGTCCACAGCTGTGCCCTGGTGGCCATGGGACATGCATGGGATCATGGGACCCCTCCCA CAACGGAGCGATGGTAGCCTGGGCTCAGGGTACGGAACTAGTCTCTCTCAGCCAACAG ACACTCGCTCATGGTGGGCCCACCGTGAAGATGGCGTGGCTCTGAGAGGCAGCCATT TCTCCTGCCAAACCAGGTTCCGGTCCCACAACCTCCGGTCCAGTCTGCAACTTCCCCTGA CTTGAAACCCACCCCAAGACCCCTACAGAGGGATGCCACCAGGCCTCAGGGCCAGAGCGT GTCTCTGGTAGCTCTGAGATCAAATCCGATGACGAGGGCGATGAGAACCTGCAAGACAC AAAATCTTCTGAGGACAAGAAATTAGATGACGACAAGAAGGATATCAAATCAATTACTAG GTCAAGATCTAGCAATAACGATGATGAGGACCTGACCCAGAGCAGAAGGCTGAGCGCGA GAAGGAACGGAGGATGGCAATAATGCCGTGAGCGCCTGAGGGTCCGAGATATCAACGA GGCTTCAGGAGCTTGGCGTATGGTCAGCTCACCTGAAGAGCGACAAGCCCCAGAC CAAGCTCCTGATTCTCCACCAGGCCGTGGCTGTACATCCTCAGCCTGGAGCAGCAAGTCG AGAAAGGAATCTGAACCCGAAAGCTGCCGTGCTGAAAAGAAGGGAGGAAGAGAAGGTGTC CTCAGAGCCTCCCCCACTCTCCTGGCTGCCACACCCCTGGGATGGGAGACGCAGCGA TCACATGGGACAGATGTGAAAAGGTTCAAGTTGCTACCTGCTTCAATTAAACAAGAGACC ACTTCCCTAACAGCTGTATTACCTAACCCACATAAACACTGCTCCTAACCCCGTTT TTTTGTAATATAAGACAAGTCTGAGTAGTTATGAATCGCAGACGCAAGAGGTTTCAGCA TTCCCAATTATCAAAAAACAGAAAAACAAAAACAAAAAATGAATGAAAGAAAGAAAGAAAG AAAAAAATGCAACTTGAGGGACGACTCTTAAACATATCACTCTGAATGTGCGACGGTAT GTACAGGCTGAGACACAGCCCAGAGACTGAATGGCAATCCTCACACTGTGGAGCAATGC ATTGTCCTAAACTCTTGGAAAAAAATATAATTAAATTGTAAGTCTGAAAAAA ATATTAAATTAAAAAAATTGAAACTTGCAATAATGAAAAAGTGTACTTCTGAAGAAA ACGACATGAACGTTTGTGTTATTACGTCAGCTAGTGTGTTCTAATTACCGGATATTG AATAGGGGAAGCCCGCTGCCCTCGTAACAAAACCAGCAAACGCTGTGGCAACGAAG TGATGACATTAGCATTCTTAGGGTAGGAGGGACAGATGGATGTTAGACCTATGACA AATATATATATAATATATAATATAATTAAAAATTAGTGAATGGTAAGCT GTGATGTCAGCTTCTCCTGTAAGGAAATTAGTACTGATAACTTTAAAGAAAGATT CTGAAATATGGATTGTTGTCTGATTGTCCTCCCTCCCCCGGTTGTTATCGTA ACCTGTAGTCCAACCTGCTTCCGGAGGGCAGTGCAGGACGAAATGCTGACCTGAAAG TTGCTCTCATCACAAATAGTAAAGTTGTTCTCAGTCTTGGGAACACAGGACT TAAAGTCACATCATGTGAGGAATTACATGCAGCATTGCCGGCGAGGAAAAAGCGT TTGCTGGCTTGCGCTGCCCTGTTACCCCTCCCCTGGGATTTAGAGGTACACGGT TAGAATGCTACAATGTTACCACTGTGCCCTCCAATGTTATATCATCGAAACATAACAT AATCAAAGTGGCTGTGATTAACAAAAAAACGATTCAAGTGTACCTACCTGTGAGCC GAAGTAGTGTGCACTGACCGAGACGTTCAAGAATACATGGTCAGATTGTTGGAAAAAA ATACAAAAATTAA
S000050	F11	138	CTGTCATTCATCAAGTCTGAAATATCGAAATGGATTAGAGAAAAATTACCCACTC CTCGGACCATCAGGACAGGGACATGGAGGGAGTGAATCAGCTGGGGGGTTTTGTGAATG GACGGCCACTCCAGATGTAGTCCGCCAAAGGATAGTGGAACTTGCCTCAAGGTGTCA GGCCCTGCGACATCTCCAGGCAGCTCGGGTCAGCCATGGTGTGTCAGCAAATTCTG GCAGGTATTATGAGACAGGAAGCATCAAGCCGGGGTGATTGGAGGATCCAACCAAAGG TTGCCACTCCCAAAGTGGGGAAAAATCGCTGAGTACAAACGCCAAAACCCCTACCATGT TTGCTGGAGATCAGGGACCGGCTGTTGGCAGAGCGAGTCTGTGACAATGACACTGTG CCAGCGTCAGCTCCATCAACAGGATCATTGGACAAAAGTACAGCAGCCCCCAATCAGC CGGCCAGCTCCAGTCACAGCATAGTGTACAGGCTCGTGACCGAGGTGTACATCGG TGAGCACCGACTCCGCGGGCTCTCATACTCCATCAGTGGCATCCTGGCATCACGTC

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			CCAGTGCGCACCCAACAAACGCAAGAGGGATGAAGGTATTCAAGGAGTCTCCAGTGCCGA ATGGCCACTCACTCCGGGCCGGACTTCCTCCGGAACGAGATGCCGGAGACCTGTTCA CACAGCAGCAGCTGGAGGTGCTGGACCGCGTGTGAGAGACAGCACTACTCTGACATCT TCACCACCAACGGAACCCATCAAGCCAGAACAGACCAACAGAGTATTCAGCCATGGCTTCAC TGGCTGGAGGCCTGGATGACATGAAGCCAACTTGACGAGCCCCACCCCCGCTGACATCG GGAGCAGCGTCCAGGCCACAGTCCTACCCATTGTCACAGGCCAGACTTGGCGAGCA CAACCCCTCCGGGGTACCTCCACACGTCCCCCGCTGGACAGGGCAGCTACTCTGCAC CGACGCTGACAGGGATGGTGCCTGGGAGTGAATTTCAGGAAAGTCCCTACAGCCACCCCTC AGTATTCTCCTACAATGATTCTGGAGGTTCCCCAACCCAGGGCTGCTTGGCTCCCCAT ACTATTACAGCCCTGCAGCCCAGGGAGCGGCCACCGGCCAGCCACTGCGTACGACC GCCACTGA
S000056	F12	139	GTTGAGCGCGAAGCAGCCGAGATGGAAGGAAGCCCTACCAACGCCACTCGGGTGGAAAGGA AAAGTCCCCTCTCCGGAGAGAGGGGACGGATCTTCACCCAGCCTGAAGCAATGGATGCC AAGCCAGCCCCCTGCTGCCAACGCCCTCTACCGGATCTGATGCTGGAGCTCTACGGAT TCGCGATGTCACAGATAGCCAGAGCGATGCCGGAGAACAGGGACAGCCCCAGGAAC CCTTCAGATCTCAGTCGGATCCTGAAGAACTCGAAGAACCCCCAGCTGTCCGCGCCGAT CCTGACGGAGGGCAGCCCCAGTCGCCCCAGCCACTCCTGCCAGTCCGAGTCTGAAGGC AGCAGAGATCCAGCCGCCAGCCAGCCTCCGAGGCAGTCCCTGCCACCGCCAGTCT GCCTCCGGGGCAGCCCCCTGTCACCCAGGTGGAGGCCAGCCGCCAGTCTGCCACC CTGGCGAGCCTGCCGCCGGCAGCCCCATCACCCCCAAGGAGCCCAGTACCCGGGCA GTCCCCCTGCTAGAGCCCATCCGCCGCTGGAGCAGTCCCTGGCCGGGAGCAATGTCA GCCTCTGCTAGGGCAGCTGCCGCTAGGGCAGCCTATGCAGGTCCACTGGTCTGGGAGCC AGGTCACTCTAGCTACTCCGCCGCTGGGATCCCTCTGCCGCCAGCAGCTGCC GCCCGGGCAGCCTCTGCTGCCGCCAGTCGCTGGGAGCAGTCCCTGCCGCCAGCAGCTGCC AGCAGGGCCCCTTAAGACCCCCAGCCCCAGATCCAGGTTGCTGACCCGCCACTCCG CGGCCCTCCGCCGCCAGTCGCTGGGAGCAGTCGCTGCCAGCAGCTGCCAGCAGCTGCC AGGTACGAGGCATCGTCTGGCATCTCGAGATCGAGTCCCTCAGCCCCGGAGGCCAAGGATCCT GGGGCCACCGGCTGCTTCAGTGGCTCTGCCGGAAACCGCCGCCCTGGCCTGCCCG AGCCACACGGTGGAGCAACCCAGTCGCAACTTCTCACCCGAGCCTCGGAAGCTGC TTGGTCTATCCGAGTGTACCGATCACGATCCCTCAGCCCCGGAGGCCAAGGATCCT ATGGAGGAGAGGCAGCAAACAGATGCGCAAAGAACGCCATTGAGATGCGAGAGCAGAAGC GCAGATAAGAAACGCAAGCTCATCGACAAGCAACTGGAGGAGGAGAACGACTAC ATGTGTACACACCGCCTGCTGCTCTAGGTGCTGGAGAGTCTGGAAAAGCACCATTTGT AAGCAGATGAGGATCCTGCATGTTAATGGGTTAACGGAGATAGTGAGAACGGCCACTAAA GTGCAGGACATCAAAACACCTGAAGGAGGCCATTGAAACCATTGTGGCCGCCATGAGC AACCTGGTCCCCCTGTTGGAGCTGCCAACCTGAGAACCCAGTTCAAGACTGGACTACATT CTGAGCGTGTGAAACGTGCCGAACCTGACTTCCACCTGAATTCTATGAGCATGCCAG GCTCTGTGGAGGATGAGGGAGTGCCTGCCGCTGCTACGAGCGCTCCAAATGAGTACAGCTG ATTGACTGTGCCAGTACTTCCTGGACAAGATTGATGTGATCAAGCAGGCCACTACGTG CCAAGTGACCAGGACCTGCTCGCTGCCGTGACCTCTGGAATCTTGAGACCAAG TTCCAGGTGGACAAAGTCACCTCCACATGTTGATGTGGCGGCCAGCGCGATGAGCG CGCAAGTGGATCCAGTGCTCAATGATGTGACTGCCATCATCTCGTGGTGGCCAGCAGC AGCTACAACATGGTATTGGGAGGACAACCAAGACTAACCGCCTGCAGGAGGCTCTGAAC CTCTCAAGAGCATCTGGAACAACAGATGGCTGCGCACCCTCTGTGATTCTCTCCTC AACAAAGCAAGACCTGCTGCTGAGAAAGTCCTCGCTGGCAAATCGAAGATTGAGGACTAC

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			TTTCCAGAGTCGCTCGCTACACCCTGAGGGATGCGACTCCCAGGCCGGGAGAGGAC CCACCGCGTACCCGGGCAAGTACTTCATTGGGATGAGTTCTGAGAATCAGCACTGCT AGTGGAGATGGCGCCACTACTGCTACCCCTACCTTGCGCCGTGGACACTGAGAAC ATCCGCCGTGCTTCAACGACTGCCGTGACATCATCCAGCGCATGCATCTCCGCAAAAC GAGCTGCTAAGAAGGAACACCCAATTAAATTCAAGCCTTAAGCACAATTAAATTAAAGA GTGAAACGTAATTGTACAAGCAGTTGGTACCCACCATAGGGATGATCAACACCGCAAC CTTCCCTTTCCCCAGTGATTCTGAAAAACCCCTCTCCCTCAGCTGCTTAGATGT TCCAAATTAGTAAAGCTTAAAGCGCCCTACAGAAGAAAAAGAAAAAGGCCACAAAAG TCCCTCTCACTTCAAGTAAATAAAAGCAGCAACAGAAATAAGAAATAATGAA ATTCAAATGAAATAATATTGTGTTGCAGCATTAAAAATCAATAAAATCAAAT GAGCAAAAAAAAAAA
S000058	F13	140	TGGACTGGGTGCGGCCGGCTGCAAGACTCTAGTCGCGCCCACGTGGCTGGGGCGGG CTGCCGTGGCGCCTAGTGATTACGTAGCGGGTGGGCCCCAAGTGCCGCTCCCTGGCGGG GCTGTTCATGGCGGTTTGGGGTCTCCAACAGCTCAGGTTGAAGTCAGCTCCCGA GGCGGGCTGCGGAGTTGAGGTTTGGCTGGTGAAATGACTGAGTACAAACTGGTGGT GGTTGGAGCAGGTGGTGGGGAAAAGCGCCCTGACGATCCAGCTAATCCAGAACCACT TGTGGATGAATATGATCCACCATAGAGGATTCTACCGAAAGCAAGTGGTATTGATGG TGAGACCTGCTGCTGGACATACTGGACACAGCTGGACAAGAGGAGTACAGTGCATGAG AGACCACTGACATGAGGACAGGCGAAGGGTCTCTGTGATTGCCATCAATAATAGCAA ATCATTGAGATATTAAACCTCTACAGGGAGCAAATTAGCGTGTAAAGATTCTGATGA TGTCCCCATGGTGTGGTAGGCAACAAGTGTGACTTGCAACAGGACAGTTGACACAAA GCAAGCCCACGAACGGCAAGAGTTACGGAATTCCATTGAGACCTCAGCCAAGAC CCGACAGGGTGTGGAGGATGCCTTTACACACTGGTAAGGGAGATAGCCAGTACCGATT GAAAAAGCTAACAGCAGTGCAGTGGCACTCAAGGTTGATGGGTGCCCCGTGTGCT GATGTGAAGACACTTGAAGATTCTGTCATCAGAAAAGGCCACTTGAAGCTGCACTG ATGCCCTGGTTCTGACATCCCTGGAGGAGACCTGTCCTGCTCTGCATCTCAGAG AAGCTCTGCTTCTGCTTCCGACTCAGTTACTGAGCACAGCCATCTAACCTGAGACC TCTCAGAATAACTACCTCTCACTCGGCTGTGCTGACCAGAGAAATGGACCTGCTCTCC CGGTGTTCTGCTGGCTGGTTCCCTAGAAACAGACACAGCCTCAGCTGGCTTGTCC TCTGAAAAGCAGTTACATTGATGCAGAGAACCAAAGTACATGCCATTCTGTTGACAA CAGTTCTTACTCTAACAGGTAACAACGACTGCTGGTATTGGGGAAATGTGATTTCT ACTGGCCTGGTTGGGGAAAATGTGATGAAATTACTTCTCCAAAATATAAT TAGTGTGCTGATTGATTTGTAATGTGATCAGCTATATTCCATGAGCTGCTCT GTATTGATGAAACAGCAACTCTAACGCAATTAAATCCACATTGCA ACAAAGTGCCTTTCTAAAGTGCCTGTAGGGCTCATTACAGTTGTAATTGGAATA GATGTGTCAGAACGACTTGTATAGGAAAGTGACTCTGAGCCATCACCTTGAGGGAAAG GTGTATGTACCTGATGGCAGATGCTTGATGACATGAAGGATAGTTCTGCTGG GATTCTCCAGGAGAAAGATGGAACGAAACATTACAAGTAATTCTATTGGAATATCTT TAATCTTTCTTTCTTTGGTAGACTACCTATAAAATATTGGAATATCTT CTAGCTTACTGATAATCTAATAATTAGGACTGCTCCATTATAATGAATTGTTCATACCA GGAAGCCCTCATTATAGTATAGGACTGTAAGGATGGCATGTTGTTACTTTAGCT GTGATTAATGATTCTCAGACCTGCTGAGGATAGTTATTAGCAGACAGGTTATATCTT TGCTGCATAGTTCTCATGGAATATATCTATCTGATGTGGAGAGAACGTGCCCTC AGTCCCTCTCAGCATCCCTCATCTCAGCCTAGAGAACGTTGAGCATCCTAGAGGGG CTTGAACAGTTATCTCGGTTAAACCATGGTCTAATGGACCGGGCATGGTTCAAAACT

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			TGAACAAGCCAGTTAGCATCACAGAGAACAGTCCATCCATATTGCTCCCTGCCTATT TTCCTGCTTACAGACTTTGCCTGATGCCTGCTGTTAGTGCTACAAGGATAAAGCTTGTG TGGTTCTCACCAGGACTGGAAGTACCTGGTGAAGCTCTGGGTAAGCCTAGATATCTTAC ATTTTCAGACCCCTATTCTTAGCCACGTGGAAACTGAAGCCAGAGTCCATACCTCCATCT CCTTCCCCCCCCAAAAAAATTAGATTAAATGTTCTTATATAGCTTTTAAAGTATTAA AACATGTCTATAAGTTAGGCTGCCAACTAACAAAAGCTGATGTGTTGTCAAATAAAGA GGTATCCTCGCTACTCGAGAGAAGAATGTAACGCCATTGATTGTTGTCAGTGGGAGG CTTGATGTTGCCTGATAATTCAATTAGTGGGTTTGTTCACATGATAACCTAAGATG TAACTCAGCTCAGTAATTCTAATGAAAACATAAATTGGATACCTTAATTGAAAAAAGCAA ACCTAATTCCAAAATGCCATTCTCTGATCTTGTAAACCTAAATTCTGAGGTC CTTGGGATTCTTTGTTTATAACAGGATCTTGCTGTGAGTCCTAGCTGGCCTCAAAC ACAATACTCTCCTGGATCAATCTCCAAGTGCCTGGGATTACAGGCACATTCCAC ACACCTGACTGAGCTGTTCTAATGAGTTTCTTAAGCAAATTCCCACACTGAA ACTAATCAGAAGGGGAACAAACATTGCTATGCTCTGAGTGCTAACACTGGCCTATT CACATGGGTTTGCACTCTAGGCAAACACTGCTGCCCTTACAACAAGGCTCAGTC ATCTCCTGAAGCTGCTGAGACCAGCACTGGCTTGTGTTTAAATATGTCTATATG ACTGGTGGTGGATCCGTCGACCTGCA
S000065	F14	141	GCTGGTGCCTCGCCGTGGCTGCTGGTACGGTCCGGAGCGATGCTGAGCCGGGCCA GCCTCTCAGCTCCGCCCTGTGCGCTGCACAGATCTAGGGGAGCCTGACGGGACGTTGACA ACGTGGAATAGGAGCAGTATCATCCCACCATGAGGTTGGGATTTAAGAGTGGAAAGATGC CAACAGCTGTGCTCTCCCATGAGGGTGTCCCCCTTCAAGTTCTCAGAACGGATGCAGGAC TGCAGATCTGTGCTGGCAACAGCAGAGGCTATATTCCAGAGGAGTCTCCAGCCGGCTG AAAGCAAATATCTATCTAAGTGCACATGTCGCAATTGTTCTGGGTGGGACATTG GTAATCCTGGTCTGTACCCACAGNGATCTTCTACGCCGTTTAAACATAAACATTGGGTT TATTAACCAGGAAAGAACAAACAAACAAAGAACACGGGGGGGGGGCTAAGAACAT ATCCG
25	S000072	F15	TGCTCCATGCCCTTGTCTCGCTGGCCCTGGCTTGCCTTAGGCTTTCTCCGCCT CTAAGTTCTTGTCCCGTCCCTAGGTCTTGTCCAGGGGGTGGGGGGGGGGACTAAG GCTGGCCTGCACACTCCAGCGAGCAGGCTATCTCTAGTTCTCGCTGCTCGGACTAGCCAT TGCCGCCGCCCTCACCTCTGCTGCAAGTAGCCTCGCCGTGGGGAGCCCTACACACGGC CGCCCTCAGCATGATGGACTTGGAGTTGCCACGCCAGACTACAGTCCAGCAGGACATG GATTGATTGACATCCTTGGAGGCAAGACATAGATCTGGAGTAAGTCGAGAAGTGT GACTTAGTCAGCGACAGAAGGACTATGAGCTGGAAAAACAGAAAAACTCGAAAAGGAA AGACAAGAGCAACTCCAGAAGGAACAGGAGAAGGCCTTTTGCTCAGTTCACTGGAT GAAGAAACAGGAGAATTCTCCCAATTGAGCCACATCCAGACAGACACCAGT GGATCCGCCAGCTACTCCAGGTTGCCACATTCCAAACAAGATGCCCTGTACTTGAA GACTGTATGCAGCTTGGCAGAGACATTCCATTGTAGATGACCATGAGTCGTTGCC CTGGATATCCCCAGCCACGCTGAAAGTTAGTCTTCACTGCCCTCATCAGGCCAGTCC CTCAATAGCTCTGGAGGCAGCCATGACTGATTAAGCAGCATAGAGCAGGACATGGAG CAAGTTGGCAGGAGCTATTCCATTCCGAATTACAGTGTCTTAATACCGAAAACAAG CAGCTGGCTGATACTACCGCTGTTCCCAGCCAGAACGCCACACTGACAGAAATGGACAGC AATTACCATTTACTCATCGATCTCTCGCTGGAAAAAGAAGTGGCAACTGTGGTCCA CATTCCCTCATGGTTTGGAGGATTCTTCAAGCAGCATCCTCTCCACTGATGATGCCAGC CAGCTGACCTCCTTAGACTCAAATCCCACCTAAACACAGATTGGCGATGAATT TCTGCTTCATAGCAGAGCCCCAGTGACGGTGGCAGCATGCCCTCCGCTGCCCATCAGT

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SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
			CAGTCACTCTCTGAACCTCTGGACGGGACTATTGAAGGCTGTGACCTGTCACTGTGTAAA GCTTCAACCCGAAGCACGCTGAAGGCACAATGGAATTCAATGACTCTGACTCTGGCATT TCACTGAACACGAGTCCCAGCGAGCGTCCCCAGAGCACCTCGTGGAGTCTCCATTAC GGAGACCCACCGCCTGGGTTCACTGACTCGAAATGGAGGAGCTAGATAGTGCCCCCTGGA AGTGTCAAACAGAACGGCCCTAAAGCACAGCCAGCACATTCTCCTGGAGACACAGTACAG CCTCTGTCAACCAGCTCAAGGGCACAGTGTCTTATGCGTAATCCAAATGTGAAAATACA ACAAAAAAAGAAGTCCCGTAGTCTGGCATCAAAAGCCCCATTACAAAAGACAAA CATTCAAGCCGCTTAGAGGCTCATCTCACAGAGATGAGCTTAGGGCAAAAGCTCCAT ATTCCATTCCCTGTGCAAAAAATCATTAACCTCCCTGTTGATGACTCAATGAAATGATG TCCAAGGAGCAATTCAATGAAGCTCAGCTCGATTGATCCGAGATAACGAGGAGAGGT AAGAATAAAAGTCGCCGCCAGAACACTGTAGGAAAAGGAAGCTGGAGAACATTGTGAGCTG GAGCAAGACTTGGGCCACTAAAGACGAGAGAGAAAACACTCAGAGAAAAGGGAGAA AACGACAGAAACCTCCATCTACTGAAAAGCGGCTCAGCACCTGTATCTGAAGTCTC AGCATGTTACGTGATGAGGATGGAAGCCTTACTCTCCAGTGAATACTCTGCAGCAA ACCAAGAGATGGAATGTGTTCTGTCCAAAAGCAAGAGCCAGATACAAAGAAAAC TAGGTTGGGAGGATGGAGCCTTCTGAGCTAGTGTGTTGTACTGCTAAAACCTC CTACTGTGATGTGAAATGAGAACACTTATAAGTAACATGAGAAATTAGCAAAG CTAGTATAGCAATAATATGAAACTTACAAGCATTAAAGTCTCAATGTTGAATCAGTT CATTAACTCTCAAGTTAATTCTAGGCACCAATTGGGAGAGTTCTGTTAAGTGA AATACTACAGAACTTATTATACTGTTCTCACTGTTACAGTCAGACTTATATGACAT CTGGCTAAAAGCAAACATTGAAAACCAACGACCACTATACTTTTATATACTGTAT GAACAGGAAATGACATTATATTAAATTGTTAGCTCATAAAAATTAAAAGGAGCTAG CACTAATAAAAGAATATCATGACT
S000083	F16	143	TATATTCCGGGGCTCGCGGCCGAGGACCCCTGGGTGCGCTGCTCTCAGCTGCCGGGT CCGACTCGCCTCACTCAGCTCCCTCCTGCTCCTGAAGGGCAGCTTCGCCGACGCTTGG CGGAAAAAGAAGGGAGGGAGGGATCCTGAGTCGAGTATAAAAGAAGCTTTCGGCG TTTTTTCTGACTCGCTGTAGTAATTCCAGCGAGAGACAGAGGGAGTGAGCGGACGGTTG GAAGAGCCGTGTGAGCCCGCTCCGGGCGACCTAAGAAGCAGCTGGAGTGA GAGGGCTTCGCTCCAGCCTGCCGCCCACCTCTCCCCAACCTGCGACTGACCAACAT CAGCGCCGCAACCCCTGCCGCCGCTGGAAACCTTGCCATTGAGCGGGAGACACTT CTCACTGGAACCTACAATCTCGAGCCAGGACAGGACTCCCCAGGCTCCGGGAGGGAAT TTTGTCATTGGGACAGTGTCTGCTCTGCCCTGCCGAGTCACTCTCCTGAAAAGA GCTCCTCGAGCTGTTGAAGGCTGGATTCTTGGCGTTGGAAACCCCGAGACAGCC ACGACGATGCCCTAACGTGAACTCACCACAGGAACATGACCTGACTACGACTCC GTACAGCCCTATTTCATCTGCGACGAGGAAGAGAATTCTATCACCAGAACAGCAGAGC GAGCTGAGCCGCCGCGCCAGTGAGGATATCTGGAAGAAATTGAGCTGCTCCACC CCGCCCTGTCCCCAGCCGCCGCTCCGGCTCTGCTCTCATCCTATGTTGCGGTGCT ACGTCCCTCTCCCCAAGGAAGACGATGACGGCGGGTGGCAACTCTCCACCGCCGAT CAGCTGGAGATGATGACCGAGTTACTGGAGGAGACATGGTAACCAAGAGCTTAC GATCCTGACGACGAGACCTTCATCAAGAACATCATCATCCAGGACTGTATGGAGCGGT TTCTCAGCCGCTGCCAAGCTGGTCTGGAGAAGCTGGCCTCCTACCAAGGCTGCGCGCAA GACAGCACCAGCCTGAGCCCCGCCGCGGGCACAGCGCTGCTCCACCTCAGCCTGTAC CTGCAGGACCTCACCGCCGCCGCTCCAGTGCTGACCTCGTCCGATTCCACGGCCCT CCGCTCAACGACAGCAGCTGCCAAATCCTGACCTCGTCCGATTCCACGGCCCT CCTTCCTCGGACTCGCTGCTGTCCTCCAGTCCCTCCCCACGGSCCAGCCCTGAGCCCTA

SAGRES TAG#	REF #	SEQ ID#	MOUSE
			SEQUENCE
			<p>GTGCTGCATGAGGGAGACACCGCCCACCAACCAGCAGCGACTCTGAAGAAGAGCAAGAAGAT</p> <p>GAGGAAGAAATTGATGTTGCTGTGGAGAAGAGGCCAACAGCAAACCTCCGCACAGCCCACGGTCTCAAG</p> <p>TCGGGCTCATCTCCATCCGAGGCCACAGCAAACCTCCGCACAGCCCACGGTCTCAAG</p> <p>AGGTGCCACGTCTCCACTCACCACTACGCCGCACCCCCCTCCACAAGGAAGGAC</p> <p>TATCCAGCTGCAAGAGGGCCAAGTTGGACAGTGGCAGGGTCTGAAGCAGATCAGCAAC</p> <p>AACCGCAAGTGCTCCAGCCCCAGGTCTCAGACACGGAGAAAACGACAAGAGGCCAGA</p> <p>CACAAACGTCTTGAACGTCAGAGGAGGAACGAGCTGAAGCGCAGCTTTTGCCCTGCGT</p> <p>GACCAAGATCCCTGAATTGAAAACAAGAAAAGGCCCCAAGGTAGTGTATCCTCAAAAAAA</p> <p>GCCACCGCCTACATCCTGTCATTCAAGCAGACGAGCACAAGCTCACCTCTGAAAAGGAC</p> <p>TTATTGAGGAAACGACGAGAACAGTTGAAACACAAACTGAAACAGCTTCGAAACAGCTTCGAAACAGCTGGT</p> <p>GCATAAAACTGACCTAACTCGAGGGAGCTGGAATCTCTCGTGAAGAGTAAGGAGAACGGT</p> <p>TCCTCTGACAGAACTGATGCGCTGGAATTAAATGCATGCTCAAAGCTAACCTCACAA</p> <p>CCTTGGCTGGGGCTTGGGACTGTAAGCTTCAGCCATAATTTAACTGCCTCAAACCTAA</p> <p>ATAGTATAAAAGAACCTTTTATGCTCCCATCTTTCTTTCTTTAACAGATT</p> <p>TGTATTAATTGTTTTAAAAAAACTTAAATAAAACGTTATAACAGTTACAAAAGATTAAAGA</p> <p>GGCCTGAAATGAAATAACTTTAATAACAGTTATAACAGTTACAAAAGATTAAAGA</p> <p>CATGTACCATATTTTTTT</p>
S000087	F17	144	<p>TATATTCCGGGGGTCTGCCGCCGAGGACCCCTGGGTGCGCTGCTCTCAGCTGCCGGGT</p> <p>CCGACTCGCCTCACTCAGCTCCCTCCTGCCCTCTGAGGGCAGCTTCGCCGACGCTTGG</p> <p>CGGGAAAAAGAAGGGAGGGAGGGATCCTGAGTCGAGTATAAAAGAAGCTTTCGGGCG</p> <p>TTTTTCTGACTCGCTGTAGTAATTCAGCGAGAGACAGAGGGAGTGGAGCAGCGGGACGGTTG</p> <p>GAAGAGCCGTGTGTCAGAGCCGCTCCGGGCGACCTAAGAAGCAGCTGGAGTGA</p> <p>GAGGGCTTGCCTCCGAGCCTGCCGCCACTCTCCCCAACCTGCGACTGACCAACAT</p> <p>CAGCGGCCGCAACCCCTGCCGCCGCTGGAAACTTGGCCATTGCAAGCGGGAGACACTT</p> <p>CTCACTGGAACCTACAATCTGAGGCCAGGACAGGACTCCCCAGGCTCCGGGAGGGAA</p> <p>TTTGTCTATTGGGACAGTGTCTGCCTCTGCCGATCAGCTCTGAAAAGA</p> <p>GCTCCTCGAGCTGTTGAAGGCTGGATTCCTTGGCGTTGGAAACCCCGAGACAGCC</p> <p>ACGACGATGCCCTCAACGTGAACTCACCAACAGGAACATGACCTGACTACGACTCC</p> <p>GTACAGCCCTATTCTGCGACGAGGAAGAGAATTCTATCACCAAGCAACAGCAGAGC</p> <p>GAGCTGCAGCCGCCGCGCCAGTGGAGGATATCTGGAAGAAATTGAGCTGCTCCCACC</p> <p>CCGCCCTGCCCCGAGCCGCCCTCCGGCTCTGCTCTCCATCTGAGCTGGCGCT</p> <p>ACGCTCTCTCCCAAGGGAAGACGATGACGGCGGGTGGCAACTCTCACCGCCGAT</p> <p>CAGCTGGAGATGATGACCGAGTTACTGGAGGAGACATGGTAACCAGAGCTCATCTGC</p> <p>GATCCTGACGACGAGACCTCATCAAGAACATCATCCAGGACTGTATGAGCGGT</p> <p>TTCTCAGCCGCTGCCAAGCTGGCTCGGAGAAGCTGGCTCTACCAGGCTGCCGCAA</p> <p>GACAGCACCAAGCCTGAGCCCCGCCGGCACAGCGCTGCTCCACCTCCAGCCTGTAC</p> <p>CTGCAGGACCTCACGCCGCCGCTCCAGTGCATTGACCCCTCAGTGGCTTCCCTAC</p> <p>CCGCTCAACGACAGCAGCTGCCCAAATCTGATCTGCCGATTCACGGCTCTCT</p> <p>CCTTCCCTGGACTCGCTGCTGTCCTCCGAGTCTCCACGGGCCAGCCCTGAGCCCCCTA</p> <p>GTGCTGCATGAGGGAGACACCGCCACCACTCACAGCAGCGACTCTGAAGAAGAGCAAGAAGAT</p> <p>GAGGAAGAAATTGATGTTGCTGTGGAGAAGAGGCCAACAGCAAACCTCCGCACAGCCCACGGTCTCAAG</p> <p>TCGGGCTCATCTCCATCCGAGGCCACAGCAAACCTCCGCACAGCCCACGGTCTCAAG</p> <p>AGGTGCCACGTCTCCACTCACCAACTACGCCGCACCCCCCTCCACAAGGAAGGAC</p> <p>TATCCAGCTGCAAGAGGGCCAAGTTGGACAGTGGCAGGGTCTGAAGCAGATCAGCAAC</p> <p>AACCGCAAGTGCTCCAGCCCCAGGTCTCAGACACGGAGGAAACGACAAGAGGCCAGA</p>

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			CACAAACGTCTTGGAACGTCAAGAGGAGGAACGAGACTGAAGCGCAGCTTTTGCCCTGCGT GACCAAGATCCCTGAATTGAAAAACAACGAAAAGGCCCCAAGGTAGTAGTCATCCTCAAAAAA GCCACCGCCTACATCCTGTCATTCAAGCAGACGAGCACAGCTCACCTCTGAAAAGGAC TTATTGAGGAAACGACGAGAACAGTTGAAACACAAACTCGAACAGCTTCGAAACTCTGGT GCATAAAACTGACCTAACTCGAGGAGGAGCTGGAACTCTCTCGTAGAGAGTAAGGAGAACGGT TCCTCTGACAGAACTGATGCGCTGGAATTAAATGCATGCTCAAAGCTAACCTCACAA CCTGGCTGGGGCTTGGACTGTAAGCTTCAGCCATAATTAACTGCCTCAAACCTAA ATAGTATAAAAAGAACCTTTTATGCTCCCACCTTTTCTTTTCTTTAACAGATT TGTATTAATTGTTTTTAAAAAAATCTAAAATCTATCCAATTTCATGAACTTAA GGCCTGAAATGTAATAACTTAAATAAAAACGTTATAACAGTTACAAAAGATTAA CATGTACCATATTTTTT	
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30	S000092	F19	TTTTTTTTTGCTTTTTTTCTTCTTCTTTCTTTCTTTCTTTGAG AGTATTGGCGACGCATTGGCGCCCTCTGAGTACGCGCAGCGAACCGGAGGCT GCGGAGGCAGAGCTGCATGCTGGCGCGTGACAGGTGGCGTGAGCAAAAGGACATT TTGGGAGTATGGGTTGGACGAGGGTGGGAGAAAAGGCAAAAGGAGACCCACGTTAGA CTGAAGAGCTAAAAGGGCACGGACTTGGCTACGCCAAGACGAAGGCCAGCCTGGGAGAGG GAGTCTCTGGGACCGGGGGGGGGGGGGCTCTGAAGCTGGCTGGTTGGGGAA GGAGGGGCTCACAAACACAGTAGGGAAGTCTGTCACTGCGAAGGGACGC GG CATCCGA CTCTCCTCTGAACTCTAAAACGTTAGCTGGCCTAGTCTCGCTGGGCCNGCC CGCGCCTCCCCGGCGCCCCAG
	S000098	F20	GCCTTAAAACGTTATTTATGTCATAAGTGCTTGCA ACTATGAGCATGTCGGT GCTCCAAAAGGCCAGGA GAGGGTGC CAGATCCTCTGAAACCCAGATGAGGGTTATGAG CCGCCATGAGGATGCTGGGA CTGAA ACCCAGGCCCTTGCACAAGCAGCAAGTGCCTCTA GCGCTCAGCCACTTCTTCATCCTCAGCATGATGAACAGAGTAAAAGCCATGAACATTGA TGAAATAAAAACATGAGTCATGTTAAAAGAACTCTGGATCTAACGGTGGACAATAGGCTA TACTGTCTCATTCA TTAAAATATGCATCTTATATAATCATAGAAAAGATGGCG AGGCACAGTCACACCAAAACATTGAGAAGATTACTCATGGGCATTAGAATTGGAGTGG TTTTAGCTTCTTCCC ACTTACTCCTGTTTCA TGTCACATGAAAAGTATTAAATGCTGC CCTAAACAGAGCAACATAGTTATTAGGGGAGACTGAGGCCTAGACAAGACAGCTCTT TACACTGAATGACTGTGGACCTGACAAAGTGGTAGATGGTGTGCTGTGACTGTCCCTGCC GTGGTAGCTACATGGTCTGAAGACTCAATTGCCGTGCGAGGAGGAATCTCTGCTCGG GCATCTGACCGCT
	S000104	F21	TATATTCCGGGGGTCTGCGCGGCCAGGACCCCTGGTGCCTCTCAGCTGCCGGGT CCGACTCGCCTCACTCAGCTCCCCTCCCTGCCTCTGAAGGGCAGCTCGCCACGCTTGG CGGGAAAAGAAGGGAGGGAGGGATCCTGAGTCGCA GTATAAAAGAAGCTTTGGCG TTTTTCTGACTCGCTGTAGTAATTCAGCGAGAGACAGAGGGAGTGAGCGGACGGTTG GAAGAGCCGTGTGCA GAGCCCGCTCCGGGGC ACCTAAGAAGGCAGCTGGAGTGA GAGGGCTTGCCCTCGAGCCTGCCGCCACTCTCCCCAACCTCGCACTGACCCACAT CAGCGGCCGCAACCCCTGCCGCCGTGGAAACTTGCCTTGAGCGGGCAGACACTT CTCACTGGAACCTACAATCTGCAGGCCAGGACAGGACTCCCCCAGGCTCCGGGGAGGGAA TTTTGTCTATTGGGACAGTGTCTGCTCTGCCCCTGCGATCAGCTCTCTGAAAAGA GCTCTCGAGCTGTTGAAGGCTGGATTCTGGCTGGAAACCCCGCAGACAGCC ACGACGATGCCCTCAACGTGAACCTCACCAACAGGAACATGACCTGACTACGACTCC GTACAGCCCTATTTCATCTGCGACGAGGAAGAGAATTCTATCACCAGCAACAGCAGAGC GAGCTGCAGCCGCCCGCCAGTGA GAGGATATCTGGAAGAAATCGAGCTGCTTCCCACC CCGCCCTGTCCCCGAGCCGCCGCTCCGGGCTCTGCTCTCCATCCTATGTTGGCTCG ACGCTCTCTCCCAAGGGAAAGACGATGACGGCGGGTGGCAACTCTCCACCGCCGAT CAGCTGGAGATGATGACCGAGTTACTGGAGGGAGACATGGTGAACCAAGAGCTTCATCTGC GATCCTGACGACGAGACCTTCAAGAACATCATCCAGGACTGTATGTGGAGCGGT TTCTCAGCCGCTGCCAAGCTGGCTCGGAGAAGCTGGCCTC GACAGCACCAGCCTGAGCCCCGCCGCCGGCACAGCGCTGCTCCACCTCCAGCCTGTAC

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S000106	F22	149	TATATTCCGGGGGTCTGGCGGCCAGGACCCCTGGGTGCGCTGCTCTCAGCTGCCGGGT CCGACTCGCCTCACTCAGCTCCCTCCTGCCCTGAAGGGCAGCTCGCCGACGCTTGG CGGAAAAAGAAGGGAGGGAGGGATCCTGAGTCGAGTATAAAAGAAGCTTTCGGGCG TTTTTCTGACTCGCTGAGTAATTCCAGCGAGAGACAGAGGGAGTGGAGCAGCGGTTG GAAGAGCCGTGTGTCAGAGCCGCCCTCGGGCGACCTAAGAAGCAGCTGGAGTGA GAGGGGCTTGCCTCCGAGCCTGCCGCCACTCTCCCAACCCCTGCAGCTGACCAACAT CAGCGGCCGCAACCCCTGCCGCCGCTGGAAACTTGCCTGCAGCGGGAGACACTT CTCACTGGAACTTACAATCTGCAGGCCAGGACAGGACTCCCCAGGCTCCGGGAGGGAA TTTGCTATTGGGACAGTGTCTCTGCCCTGCCCGCATCAGCTCTCTGAAAAGA GCTCCTCGAGCTGTTGAAGGCTGGATTCCCTTGGCGTTGGAAACCCCGAGACAGCC ACGACGATGCCCTAACGTGAACCTCACCAACAGGAACATGACCTCGACTACGACTCC GTACAGCCCTATTCATCTGCAGGCCAGGAGAAGAATTCTATCACCAGCAACAGCAGAGC GAGCTGCAGCCGCCGCGCCAGTGGAGATATCTGGAGAAGAAATTGAGCTGCTCCACC CCGCCCCCTGCCCGAGCCGCCGCTCCGGCTCTGCTCTCCATCCTATGTTGGCGTGC ACGTCCTCTCCCCAAGGGAGACGATGACGGGGGGGGCAACTCTCCACCGCCGAT CAGCTGGAGATGATGACCGAGTTACTGGAGGAGACATGGTAACCAGAGCTCATCTGC GATCCTGACGAGACCTTCAAGAACATCATCATCCAGGACTGTATGTGGAGCGGT TTCTCAGCCGCTGCCAAGCTGGCTCGGAGAAGCTGGCCTCTACCAAGGCTGCCGCAA GACAGCACCAGCCTGAGCCCCGCCGCCAGCAGCTGCTCCACCTCCAGCCTGTAC CTGCAGGACCTCACCGCCGCCGTCCGAGTCATTGACCCCTCAGTGGTCTTCCCTAC CCGCTCAACGACAGCAGCTGCCAAATCCTGTACCTCGCCATTCCACGGCCCTCT CCTTCCTCGGACTCGCTGCTGCCAGTCCTCCCACGGCCAGCCCTGAGCCCCTA GTGCTGCATGAGGAGACACCGCCACCACCAAGCAGCAGTCTGAAGAAGAGCAAGAAGAT GAGGAAGAAATTGATGTGGTGTCTGTGGAGAAGAGGCAAACCCCTGCCAAGAGGTGGAG TCGGGCTCATCTCCATCCGAGGCCACAGCAAACCTCCGCACAGCCACTGGTCTCAAG

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S000107	F3	150	TATATTCCGGGGGTCTGCGCGGCCAGGACCCCTGGGTGCGCTGCTCTCAGCTGCCGGGT CCGACTCGCCTCACTCAGCTCCCCTCCTGCCTCTGAAGGGCAGCTCGCCACGCTTGG CGGGAAAAAGAAGGGAGGGAGGGATCCTGAGTCGAGTATAAAAGAAGCTTTCGGCG TTTTTCTGACTCGCTGTAGTAATTCCAGCGAGAGACAGAGGGAGTGAGCGGACGGTTG GAAGAGCCGTGTGCGAGAGCCGCGCTCCGGGGCAGCTAAGAAGGCAGCTCTGGAGTGA GAGGGCTTGCCTCCGAGCCTGCCGCCCACACTCTCCCCAACCTCGACTGACCCACAT CAGCGGCCGCAACCCCTGCCGCCGCTGGAAACTTTGCCATTGCAAGCGGGAGACACTT CTCACTGGAACTTACAATCTGCGAGCCAGGACAGGACTCCCCAGGCTCCGGGAGGGAA TTTGTCTATTGGGGACAGTGTCTCTGCCTCTGCCGCGATCAGCTCTGAAAAGA GCTCCTCGAGCTGTTGAAGGCTGGATTCTGGGGTTGGAAACCCCGAGACAGCC ACGACGATGCCCTAACGTGAACTTCACCAACAGGAACATGACCTGACTACGACTCC GTACAGCCCTATTTCATCTGCGACGAGGAAGAGAATTCTATCACCAGAACAGCAGAGC GAGCTGCAGCCGCCGCGCCAGTGGAGGATATCTGGAAGAAATTGAGCTGCTCCACC CCGCCCCCTGTCCCCGAGCCGCCGCTCCGGCTCTGCTCTCCATCCTATGTCGGGTCGCT ACGTCCTCTCCCCAAGGGAAAGACGATGACGGCGGGTGGCAACTTCTCACCGCCGAT CAGCTGGAGATGATGACCGAGTTACTGGAGGAGACATGGTAACCAGAGCTTCATCTGC GATCCTGACGACGAGACCTTCATCAAGAACATCATCATCCAGGACTGTATGGAGCGGT TTCTCAGCCGCTGCCAAGCTGGTCTCGAGAGAAGCTGGCCTCCTACCAGGCTGCCGCAA GACAGCACCAGCCTGAGCCCCGCCGCGACAGCGCTGCTCCACCTCAGCCTGTAC CTGCAGGACCTCACCGCCGCCGCTCCGAGTGCATTGACCCCTCAGTGGCTTTCCCTAC CCGCTCAACGACAGCAGCTGCCAACCTGTACCTCGCTCCGAGTCTCCACGGCTTCT CCTTCCCGGACTCGCTGCTGTCCTCCGAGTCTCCACGGGCCAGCCCTGAGCCCCCTA GTGCTGCATGAGGAGACACCGCCACCACCAAGCAGCGACTCTGAAGAAGAGCAAGAAGAT GAGGAAGAAATTGATGTTGCTGAGAGGCAAACCCCTGCCAAGAGGTCGGAG TCGGGCTCATCTCCATCCCGAGGCCACAGCAAACCTCCGACAGCCACTGGCTCTCAAG AGGTGCCACGTCTCCACTCACCAAGCACAACATCGCCGACCCCCCTCCACAAGGAAGGAC TATCCAGCTGCCAAGAGGGCCAAGTTGGACAGTGGCAGGGCTCTGAAGCAGATCAGCAAC AACCGCAAGTGTCCAGCCCCAGGTCTCAGACACGGAGGAAACGACAAGAGGCGGACA CACAACGTCTTGGAACGTCAGAGGAGGAACGAGCTGAAGCGCAGTTTTGCCCTGCGT GACCAGATCCCTGAATTGGAAAACAACGAAAAGGCCCCAAGGTAGTGTACCTCTGAAAA GCCACCGCCTACATCCTGTCCATTCAAGCAGACGAGCACAAGCTCACCTCTGAAAAGGAC

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S000113	F24	151	GGCACGAGCCGAGTTGGAGGAAGCAGCGGCAGCGGCAGCGGTAGCGGTGAGGAC GGCTGTGCAGCCAAGGAACCGGGACAGCGAACGCGACGGCAGGTCGAGCTGGATCGCAGG AGCCTGGGAGCTGGGAGCTCAGAGGCCGCTGAAGGCCAGGCTGGCAGAGGAAGGAAGC GAGCCGACCCGGAGGTGAAGCTGAGAGTGGAGCGTGGCAGTAAATCAGACGACAGATGG ACAGTGTGACAGGAACGTCAGAGAGGATTGGGCCTCGCTGCGAGAGTCAGCCTGGAGTCA AGGTGTTACAAGTTGCTGAGAAGGACACGTGGAGGACGGTGGCGCGCGAGGGAGAGC CCTGCTTCAGTCACCCCGTTGATGGAGGACAGATGGACAGCAGCCGACGGCAGTCAC CTCTCTAAACCTTGGATAGTGGCCTTGTGCTCTGGACACCTGTTGGGAGTT AGCCCATTCTCTGAACACTCTTCTCTTAAACGTAACACTCGGACGGCAGTGTGCGAGCC AGCTCCTCTGTCAGGGCACTAGAGCTGCAAGACATGAGTGCAGAGGGCTACCAGTACAG AGCACTGTACGACTACAAGAAGGAGCGAGAGGAAGACATTGACCTACACCTGGGGACAT ACTGACTGTGAATAAAGGCTCTTAGTGGCACTGGATTCACTGATGCCAGGAAGCCCG GCCTGAAGATATTGGCTGGTAAATGGCTACAAATGAAACCACTGGGAGAGGGAGACTT TCCAGGAACCTACGTTGAATAACATTGGAAGGAAAAGAATTTCACCCCTACTCCAAAGCC TCGGCCCCCTCGACCGCTCCTGTTGCTCCGGTTCTCAAAACTGAAGCTGACACGGA GCAGCAAGCAGTTGCCCTCCCTGACCTGGCCGAGCAGTTGCCCTCTGATGTTGCC GCCTCTCTTATAAGCTCTGGAAGCCATTGAGAAGAAAGGACTGGAATGTTGACTCT ATACAGAACACAAAGCTCAGCAACCCCTGCAGAATTACGACAGCCTCTGATTGTGATGC CGCGTCAGTGGACTTGGAGATGATCGACGTACACGTCTTAGCAGATGCTTCAAACGCTA TCTCGCCGACTTACCAAATCCTGTCATTCTGTAGCTGTTACAATGAGATGATGTC AGCCCAAGAAACTACAGAGCCCTGAAGACTGCATCCAGCTGTTGAAGAAGCTCATTAGATT GCCTAATATACTCATCAGTGTGGCTTACGCTTCAGTATTGCTCAAGCATTTC GCTCTCTCAAGCCTCCAGAAAAACCTTTGAATGCAAGAGTCTCTGAGATTTCAG CCCCGTGCTTTCACTGATTTCCAGGCTCAGCTGATAACTGAAACACCTCATAAAAGC GATAGAGATTAACTCAACGGAATGGAATGAGAGACAGCCAGCACCAGCACTGCC CAAACCAAGCCACTACTGTAGCCAACACAGCATGAAACAAATATGCTTGC GGATGCTGAATGGTACTGGGAGACATCTCAAGGGAAGAAGTGAATGAAAACCTCC CACTGCTGATGGGACCTTTGGTACGAGACGCATCTACTAAATGCACGGCATTACAC TCTTACACCTAGGAAAGGAGGAATAACAAATTAACTCAAATCTTACCGTGTAGGAAA ATATGGCTTCTGATCCATTAAACCTCAACTCTGTGGTTGAGTTAATAAACCACTACCG GAATGAGTCTTAGCTCACTACACCCCAAGCTGGATGTGAAGTTGCTCTACCCAGTGT CAAATACCAAGCAGGATCAAGTTGTCAAAGAAGATAATTGAGAGTGTAGGGAAAAATT ACATGAATATAACTCAATTCAAGAAAAAGTCGGAATATGATAGATTATGAGGA GTACACCCGTACTTCCCAGGAAATCAAATGAAAAGAACGGCTATCGAAGCATTAA AACCATAAAAATATTGAGAACAATGCCAAACCCAGGAGCGGTACAGCAAAGAATACAT AGAGAAGTTAAACCGCAAGGCAACGGAGAAAGAAATTCAAAGGATTATGCATAACCATGA TAAGCTGAAGTCGCGTATCAGTGAGATCATTGACAGTAGGAGGAGGTTGAGAAGACTT

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SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
			GAAGAAGCAGGCAGCTGAGTACCGAGAGATCGACAAACGCATGAACAGTATTAAGCCGGA CCTCATCCAGTTGAGAAAGACAAGAGACCAATACTTGATGTGGCTGACGCAGAAAGGTGT GCGGCAGAAGAAGCTGAACGAGTGGCTGGGAATGAAAATACCGAAGATCAATACTCCCT GGTAGAAGATGATGAGGATTGCCCCACCATGACGAGAAAGACGTGGATGTCGGGAGCAG CAACCGAAACAAAGCGGAGAACCTATTGCGAGGGAAAGCCAGACGGCACTTCCCTGTCCG GGAGAGCAGTAAGCAGGGCTGCTATGCCCTCGTAGTGGTAGACGGCGAAGTCAAGCA TTGCGTCATTAACAAGACTGCCACCGGCTATGCCCTGGCTTGGCAGGCCCTACAACCTGTACAG CTCCCTGAAGGAGCTGGTGTACATTATCAACACACCCCTCGTCGAGCACAATGACTC CCTCAATGTCAACACTAGCATAACCCAGTATATGCACAACAGAGGCAGTGAAGCCTGCCCT CGGATCCAGTCCCTCACCTCAAGCCACCCAGGCCTCTGAGAAGCAAAGGGCTCTCTC CAGCCCGACCTGTGAAGCTGAGCTGCAGAAATGAAGCCGGCTGTCTGCACATGGGACTAGA GCTTCTGGACAAAAGAAGTCGGGAAGACACGCAGCCTCGGACTGTTGGATGACCAG ACGTTCTAACCTTATCCTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTT TTCTTCTTCTTCTTCTTCTTCTTCTAATTAAAGCCACAACACACACACAGAG AGAAAGAAATGCAAAATCTCCGTGCAGGGACAAGAGGCCTTAAACCATGGTGTG TTAACGCTTCTGAGCTTACAGCTACAAGTGGACTTGGAGACAGAAGGTAGAC AGGGCGGAAGAGCCTGCCCTGGGCCGCTGGTCCAGCCTGGTAGCCTGGGTGTCGC TGGGTGTGGTAACCCAGACACATCACACTGTGGATTATTCCTTTAAAGAGCGAAT GATATGTATCAGAGAGCCCGTCTGCTCACGCAGGACACTTGAGAGAACATTGATGCAG TCTGTTGGAGGAAAATGAAACACCAGAAAACGTTTGTAAACTTCAAGTCAGC AACCAACAACCCACCAACAGAAAAA
35	S000114	F25	GTTGCCGGTTAGGGTGTGCTGTAGTGGCGATACGTCCGCCGTGTCCTGCCAGTGAGG GATCCGAGCCCGCAGCGAGTGCATGGAGGGCCAGCGCGTGGAGGAGCTGTCGCCAGGC AGAGCAGGAGGAGGCGGAGAAGCTGCAGCGCATCACGGTGACAAAGGAGCTGGAGCTGG GTTGACCTGGCAACCTGCTGGCTCGGACCGCAACCCCCCGACCGTGCTGCCAGGC CGGGCCGTCGCCGGAGGCCGAGCTGCCGGCCCTGGCGCGGGACAACACGCAGCTGCTCAT CAACCAGCTGTGGCGCTGCCGACCGAGCGCGTGGAGGAGGCGGTGGTGCAGCCTTG GGAGCCCCGCACTCGCCTGCCCGAGAAGCCGCTGCCCGACCGCCGCTCACCC CTGGCAGCAGTCGCGCCCTAAGGGATCCGTCAGAAGAACACCCCTCGTGTG GGACGAGGCTAGTGGCAGTGGCGCCGTTGGCTACAAGCGCAGGGATGACAC TAAAGAATGGCTGATCGAGGTGCTGGAGCGCCGACCCATGAAAGACCAGTTGCCAA GAGGACTCAGGCCAAGAAAGAACCGCTGGCCAAGAATGAGCTGAACCGTCTGCCAACCT GGCTCGCGCGACAAGATGCGATGCCAGCTGCCGCTGCACCCACTGGACACCA GAGTAAGGAAGAGCTGGCCGCGCCATGCAAGTGGCAAGGTTCCACCGCTCGGTGG ACGCTCCAGGAGCGCCTTCCAAGGGAGAAAGCTCCCCGGGCTCCGGCAAGAAGAGGAA GTTTCAGCCCCCTTTGGGACTTCCGAGCCGAGAAAAAGAACCGAGTTGGAGCTACTTCG AGTCATGAACAGCAAGAACCTCGGCTGGACGTGACGAGGGCCACCAAGCAGATGAG GGAAGAGGACCGAGGAGGAGGCTGCCAAGAGGAGGAAATGAGCCAGAAAGGCAAGAGGAA AGGGGGCGGCAAGGACCTCGGGCAAGAGAAGGGCGGCCGGGTAGGGAGGAA GAGGAAAGGAGGCTGGGAAGCAAAAAGCATTCCTGGCCTTCTGCTTAGCTGCCAGAA GAAAGGAGTGGCGCCCAAGGTGGGAAGAGGAGGAAAGTAGCGTTCCCCTCGGGACCAG TTCTGAAAAGCTGGGACTGTACTAAAGTTAACCTGGCGGTAGGTGGCGCTGCCCT CAGTGACATTGACATTAAGGACGGGTTGCCCTCCCTCGAGTCAGTGTGGACGAGT TAATAGAGACACTGACTGAAATTGGTGTATTGAGAATTATAGAAATGATATAGCCAG AACCAAGAATAAGTTAAGGCCCTGCCCTTATCTGACTTTGGATACTGCCAGTAG

SAGRES TAG#	REF #	SEQ ID#	MOUSE
			SEQUENCE
			ATGGTTCAACATTTGCATTATTTATAACAAAGCTGTGTTTATCAAAGCGGG GAGGGCGGGAAAAATTATCTACCTGTGATTGCAAGTATTGAAATGGATGCAGGTA CCTGGTGTGCTTTAACTTACTGTCGGTAGAGGTTGCATGTGAAGCCAGTAACCTGG GCACCAATATGGAGTGTGCTTGAGAAAACAAAGTAGTTACAGTGGTTCTAAAAAAGACC CCTGTTAGAAAACTTGGCCCTAACTATAATATTAAAAGTATAGTGTGTTGGTG TTGGTCAGGTGGTGCATTGCCAATGGATTGCTTAAGTCCAGAAATAGTGTCACTT TGTTGTAACCGGTGGCTTGTTAATTGGCTGGGTTAGATATTGTCAAAATATCT GGCATTCACTATGGAACCAAGGCTGCCCTGGAACTCAGGCCAAGTGTGAGATTATAAT CGAGCAGCAGATTCATGTTATTCCTGCTTAGATGTTTCCCTGTTATTGTCCTA TTTGTCTTAATAAACTATCTTGCATAAAAAAAAAAGGCCACA
S000116	F26	153	TATATTCCGGGGTCTGCGCGGCCGAGGACCCCTGGGTGCGCTGCTCTCAGCTGCCGGG CCGACTCGCCTCACTCAGCTCCCTCCTGCCCTGAAGGGCAGCTCGCCGACGCTGG CGGAAAAAGAAGGGAGGGAGGGATCCTGAGTCGCAGTATAAAAGAAGCTTTCGGCG TTTTTCTGACTCGCTGTAGTAATTCCAGCGAGAGACAGAGGAGTGAAGCGGACGGTTG GAAGAGCCGTGTGTCAGAGCCGCGCTCCGGGCGACCTAAGAAGCAGCTCTGGAGTGA GAGGGCTTCCTCCGAGCCTGCCGCCACTCTCCCCAACCTGCACTGACCCAACAT CAGCGGCCGCAACCCCTGCCGCCGCTGGAAACTTGCCTATTGCAAGCGGGCAGACACTT CTCACTGGAACTTACAATCTGCGAGCCAGGACAGGACTCCCCAGGCTCCGGGAGGGAT TTTGTCTATTTGGGGACAGTGTCTGCCTCTGCCGCTGCCCCTGAGCTCTCTGAAAAGA GCTCCTCGAGCTGTTGAAGGCTGGATTCTTGGCTGGAAACCCCGCAGACAGCC ACGACGATGCCCTCAACGTGAACCTACCAACAGGAACTATGACCTGACTACGACTCC GTACAGCCCTATTCATCTGCGACGAGGAAGAGAATTCTATCACAGCAACAGCAGAGC GAGCTGAGCCGCCGCGCCAGTGAAGGATATCTGGAAGAAATTGAGCTGCTCCACC CCGCCCCCTGTCCCCGAGCCGCCGCTCCGGCTCTGCTCTCCATCCTATGTTGCCGCTGCT ACGTCTTCTCCCCAAGGGAAGACGATGACGGGGCGTGGCAACTCTCCACCGCCGAT CAGCTGGAGATGATGACCGAGTTACTGGAGGAGACATGGTAACCAGAGCTCATCTG GATCCTGACGACGAGACCTTCAAGAACATCATCATCCAGGACTGTATGGAGCGGT TTCTCAGCCGCTGCCAAGCTGGTCTGGAGAAGCTGGCTCCTACCAAGGCTGCCGCAA GACAGCACCAGCCTGAGCCCGCCGCCGACAGCGCTGCTCCACCTCAGCCTGTAC CTGCAAGGACCTCACCGCCGCCGCTCGAGTGCATTGACCCCTCAGTGGCTTCCCTAC CCGCTCAACGACAGCAGCTGCCAAATCCTGTACCTGCTCCGATTCCACGGCTTCT CCTTCCCTGGACTCGCTGCTGCCCTCGAGTCCCTCCCACGGGCCAGCCCTGAGCCCTA GTGCTGCATGAGGAGACACCGCCACCCACCAGCAGCGACTCTGAAGAAGAGCAAGAAGAT GAGGAAGAAATTGATGTGGTGTGAGAAGAGGCAAACCCCTGCCAAGAGGTCGGAG TCGGGCTCATCTCCATCCGAGGCCACAGCAAACCTCCGACAGCCACTGGCTCTCAAG AGGTGCCACGTCTCCACTCACCAGCACAACACTACGCCGACCCCCCTCCACAAGGAAGGAC TATCCAGCTGCCAAGAGGGCAAGTTGGACAGTGGCAGGGCTCTGAAGCAGATCAGCAAC AACCGCAAGTGTCCAGCCCCAGGTCTCAGACACGGAGGAAACGACAAGAGGCGGACA CACAAACGTCTGGAACGTCAGAGGAGGAACGAGCTGAAGCGCAGCTTTTGCCTGCGT GACCAAGATCCCTGAATTGGAAAACAACGAAAAGGCCCCAAGGTAGTGATCCTCAAAAAA GCCACCGCCTACATCTGTCATTCAAGCAGACGAGCACAAGCTCACCTCTGAAAAGGAC TTATTGAGGAAACGACGAGAACAGTTGAAACACAAACTCGAACAGCTCGAAACTCTGGT GCATAAAACTGACCTAACTCGAGGAGGAGCTGGAATCTCTCGTGAGAGTAAGGAGAACGGT TCCTCTGACAGAACTGATGCGCTGGAATTAAATGCATGCTCAAGCCTAACCTCACAA CCTGGCTGGGCTTGGACTGTAAGCTCAGCCATAATTAACTGCCTCAAACCTTAA

MOUSE			
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
			ATAGTATAAAAGAACCTTTTATGCTTCCCATCTTTTCTTTCTTTAACAGATT TGTATTAATTGTTTTAAAAAAATCTAAAATCTATCCAATTCCCAGTAAATAG GGCCTGAAATGTAATAACTTTAATAAAAACGTTATAACAGTTACAAAAGATTAAAGA CATGTACCATAATTTTTTT
S000118	F27	154	TATATTCCGGGGTCTGC CGGCC GAGGACCCCTGGTGCGCTGCTCTCAGCTGCCGGT CCGACTCGCCTCACTCAGCTCCCCTCCTGCCTCTGAAGGGCAGCTCGCCGACGCTTGG CGGAAAAAGAAGGGAGGGAGGGATCCTGAGTCGCACTGATAAAAGAAGCTTTCGGCG TTTTTCTGACTCGCTGTAGTAATTCCAGCGAGAGACAGAGGGAGTGAGCGGACGGTTG GAAGAGCCGTGTGAGAGCCCGCTCCGGGCGACCTAAGAAGGCAGCTGGAGTGA GAGGGCTTGCCTCCGAGCTGCCGCCCCACTCTCCCCAACCTGCGACTGACCAACAT CAGCGGCCGCAACCCCTGCCGCCGCTGGAAACTTTGCCATTGCAAGCGGGCAGACACTT CTCACTGGAACTACAATCTGCAGGCCAGGACTCCCCAGGCTCCGGGAGGAAT TTTGTCTATTGGGGACAGTGTCTGCCTCTGCCGATCAGCTCTCTGAAAAGA GCTCCTCGAGCTGTTGAAGGCTGGATTCCCTGGCGTTGGAAACCCCGCAGACAGCC ACGACGATGCCCTAACGTGAACTCACCAACAGGAACATGACCTGACTACGACTCC GTACAGCCCTATTCATCTGCAGGCCAGGAGAATTCTATCACAGCAACAGCAGAGC GAGCTGCAGCCGCCGCGCCAGTGGAGATCTGGAGAAATTGAGCTGCTTCCACC CCGCCCCCTGTCCCCGAGCCGCCGCTCCGGCTCTGCTCTCCATCCTATGTCGGTGC ACGTCCCTCCCCAAGGGAAGACGATGACGGCGGGTGGCAACTCTCACCGCCGAT CAGCTGGAGATGATGACCGAGTTACTGGAGGAGACATGGTAACCAGAGCTTCT GATCCTGACGAGACCTTCAAGAACATCATCCAGGACTGTATGGAGCGGT TTCTCAGCCGCTGCCAAGCTGGTCTGGAGAAGCTGGCCTCTACCAGGCTGCCGAA GACAGCACCGCCTGAGCCCGCCGCCGCTCGAGTGCATTGACCCCTCAGGGTCT CTGCAGGACCTCACGCCGCCGCGTCCAGTGCATTGACCCCTCAGGGTCT CCGCTAACGACAGCAGCTGCCCAAATCCTGTACCTCGCTCCAGGCTCACCGCT CCTTCCCTGGACTCGCTGCTGTCCCTGGAGTCTCTCCACGGGCCAGCCCTGAG GTGCTGCATGAGGAGACACGCCACCACCGAGCAGCAGCTGAAGAAGAGCAAGA GAGGAAGAAATTGATGTTGGTCTGGAGAAGAGCAACCCCTGCCAAGAGGCTGG TCGGGCTCATCCATCCGAGGCCACAGCAAACCTCCGACAGCCACTGGTCTCAAG AGGTGCCACGCTCCACTCACAGCACAACGAGCTGCCGACCCCCCTCCACAAG TATCCAGCTGCCAAGAGGGCAAGTTGGACAGTGGCAGGGCTCTGAAGCAGATCAG AACCGCAAGTGTCCAGCCCAAGGTCTCAGACACGGAGGAAACGACAAGAGGCG CACAAACGTCTGGAACGTCAGAGGAGGAACGAGCTGAAGCGCAGCTTTTGCC GACCAAGATCCCTGAATTGGAAAACAACGAAAAGGCCAGGTAGTGTATCT GCCACCGCCTACATCCTGTCCATTCAAGCAGACGAGCACAAGCTCACCTCT TTATTGAGGAAACGACGAGAACAGTTGAAACACAAACTCGAACAGCTTC GCATAAAACTGACCTAACTCGAGGAGGAGCTGGAACTCTCGTGAGAGTAAG GGTCTGACAGAACTGATGCGCTGAAATTAAATGCATGCTCAAAGCTAAC CCTGGCTGGGCTTGGACTGTAAGCTTCAGCCATAATTAACTGCCTCAA ATAGTATAAAAGAACCTTTTATGCTTCCCATCTTTCTTTCTTTAACAGATT TGTATTAATTGTTTTAAAAAAATCTAAAATCTATCCAATTCCCAGTAAATAG GGCCTGAAATGTAATAACTTTAATAAAAACGTTATAACAGTTACAAAAGATTAAAGA CATGTACCATAATTTTTTT
S000121	F28	155	TATATTCCGGGGTCTGC CGGCC GAGGACCCCTGGTGCGCTGCTCTCAGCTGCCGGT CCGACTCGCCTCACTCAGCTCCCCTCCTGCCTCTGAAGGGCAGCTCGCCGACGCTTGG

			MOUSE
SGRES TAG#	REF #	SEQ ID#	SEQUENCE
			CGGGAAAAAGAAGGGAGGGAGGGATCCTGAGTCGCACTATAAAAGAAGCTTTGGGG TTTTTCTGACTCGCTGTAGTAATTCCAGCGAGAGACAGAGGGAGTGAGCGGACGGTTG GAAGAGCCGTGTGAGAGGCCGCTCCGGGGCGACCTAAGAAGGCAGCTCTGGAGTGA GAGGGCTTGCCCTCGAGCCTGCCGCCACTCTCCCCAACCTGCGACTGACCAACAT CAGCGGCCGCAACCCTGCCGCCGCTGGAAACTTGGCCATTGAGCAGGGAGACACTT CTCACTGGAACCTACAATCTGCAGGCCAGGACAGGACTCCCCAGGCTCCGGGAGGGAA TTTGTCTATTGGGGACAGTGTCTGCCTCTGCCCTGCCGATCAGCTCTGAAAAGA GCTCCTCGAGCTGTTGAAGGCTGGATTTCTTGGCGTTGGAAACCCCGCAGACAGCC ACGACGATGCCCTAACGTGAACTCACCAACAGGAACATGACCTCGACTACGACTCC GTACAGCCCTATTCATCTGCAGCAGGAAGAGAATTCTATCACCAACAGCAGAGC GAGCTGCAGCCGCCGCGCCAGTGAGGATATCTGGAAGAAATTGAGCTGCTCCCACC CCGCCCTGTCCCCGAGCCGCCGCTCCGGGCTCTGCTCTCCATCCTATGTTGCCGCT ACGTCTTCTCCCCAAGGGAAGACGATGACGGCGGGTGGCAACTTCCACCCGCGAT CAGCTGGAGATGATGACCGAGTTACTGGAGGAGACATGGTGAACCAAGAGCTCATCTG GATCCTGACGAGACCTCATCAAGAACATCATCATCCAGGACTGTATGGAGCGGT TTCTCAGCCGCTGCCAAGCTGGTCTCGGAGAAGCTGGCCTCCTACCAGGCTGCCGCAA GACAGCACCAGCCTGAGCCCCGCCGCGGACAGCGCTGCTCCACCTCAGCCTGTAC CTGCAGGACCTCACGCCGCCGCTCCAGTGATTGACCCCTCAGGGTCTTCCCTAC CCGCTCAACGACAGCAGCTGCCAAATCCTGTACCTCGCCATTCCACGGCTTCT CCTCTCGGACTCGCTGTCCTCGAGTCTCCACGGCCAGCGACTCTGAAGAAGAGCAAGA GTGCTGCATGAGGAGACACCGCCACCCACCAGCAGCGACTCTGAAGAAGAGCAAGA GAGGAAGAAATTGATGTTGAGGAGACAGGCAACCTCCGACAGCCACTGGTCTCAAG AGGTGCCACGTCTCCACTCACCGACAACACTACGCCGACCCCCCTCCACAAGGAAGGAC TATCCAGCTCCAAGAGGGCAAGTTGGACAGTGGCAGGGCTCTGAAGCAGATCAGCAAC AACCGCAAGTGTCCAGCCCCAGGTCTCAGACACGGAGGAAACGACAAGAGCGGACA CACACGTCTGGAACGTCAAGAGGAGGAACGAGCTGAAGCGCAGCTTTTGCCCTGCGT GACCAAGATCCCTGAATTGAAAACAACGAAAAGGCCCCAAGGTAGTGTATCCTCAAAAAA GCCACCGCTACATCCTGTCATTCAAGCAGACGAGCACAGCTCACCTCTGAAAGGAC TTATTGAGGAAACGACGAGAACAGTTGAAACACAAACTCGAACAGCTTCGAAACTCTGGT GCATAAACTGACCTAACTCGAGGAGGAGCTGGAAATCTCTCGTGAGAGTAAGGAGAACGGT TCCTCTGACAGAACTGATGCGCTGGAATTAAATGCATGCTCAAAGCTAACCTCACAA CCTGGCTGGGCTTGGACTGTAAGCTTCAGCCATAATTAACTGCCTCAAACCTAA ATAGTATAAAAGAACCTTTTATGCTCCATCTTTCTTTCTTTAACAGATT TGTATTAATTGTTTTAAAAAAATCTAAAATCTCAATTCCCAGTAAACAGTTATAACAGTAAAG GGCCTGAAATGTAATAACTTAATAAAAACGTTATAACAGTTACAAAAGATTAAAGA CATGTACCATATAATTTTTT

Contigs assembled from the human EST database by the NCBI having homology with all or parts of the LA
40 nucleic acid sequences of the invention are depicted in Table 3.

TABLE 3

			HUMAN
SGRES	REF	SEQ	SEQUENCE

TAG#	#	ID#	
S000010	F29	156	<p>GTGTGGCTGGACCTCGTGCAGCTGCCATTGCCAGTGGATGGAAGAAGAAAGGGCT CCGCGCAAGCGCCGATGGCGCGCCTCCAGTGCCTGGCAGCGACTCGGAGGACGCG CGAGTTGCAGATCCATGTGCTGGACAGATGACTGCCCTGGGCCAGCTGGGACCTG GAAGACCCCTGCCAACCTCCCCACCTCGGAATGCACCTCGCGATGTGGAGCCCGACAC CGGGCAGATGGCTGCCTGCCAGAACAGCAAGACAGAAGAACGCTGGCAGGCTTCCA GTCCATGGGCCCTGAGCTACCCGGTGTCAAAGGCATCATGACACGAAGGGTACAAGGT GCCAACACCCATCCAGAGGAAGACCATCCGGTATCTGGATGGCAAGGACGTGGTGGC CATGGCCCGGACGGGAGTGGCAAGACATGCTGCTCCCTCCCATAATGCCAGCGGCT CAAGACCCACAGTTGCCAGACCCGGGCCCTGTGCCCTCATCCTCTGCCGACCCGAG AGCTGGCCCTTGCAAGACCCCTGAAGTTCACTACGGAGCTAGGCCAGTCCCTGGCCTCAAG ACTGCCCTGATCTGGTGGCGCCCGGATGCCACCCGCCCTCGCAGCCCTGCACCGCAA ATCCCGACATACCTTGGCAGGCCGGGACCGTTGGGCTGTGGCAATTGAGC CTGCAGCTCCAGTTGGCCTCCGTGGTGGCCGACCCCTGCCGCGCTTCGCC GCGTTCTCGCTCATCCCCCTCCGTGGCCTTCCGCCGCCCTCCCCGGGGGGGGGGGGGGGG ACCGGCGGGCGCTCCCTGCGCCGCCCTCCCCACCCCTGTCGTGCTGGCGATTGCCCCGG CTGTGCCCTCCCCGGGGGGGGGGTCAACCCGGTGCGGGGACTACACCCCTCGCGCCTCA GTGCCCTCTCCCCGGGGAGGACCCACGCCGCTGCC</p>
S000013	F30	157	<p>CACACCGCAGTATGCCGTGCCCTTAACCTCTGAGCTGCGCAGCCGGCCGGCGCTGGT TGAACAGACTGCCGTGACTGGCGTGGCCTGGAGGGACTCAGCAAATTCTCCCTGCC AACTTGGCAACAGTTGCCCTGGGTAGCTCTACACAACTCTGTCCAGCCCACAGCAATGAT TCCAGAGGCCATGGGAGTGGACAGCAGCTAGCTGACTGGAGGAATGCCACTCTCATGG CAACCACTACAGCACTATCATGCAGCAGCCATCCTGCTGACTAACCATGTGACATTGGC CACTGCTCAGCCTCTGAATGTTGGTGTGCCATGTTGTCAGACAACAAATCCAGTT CCTCCCTCGAAGAAGATAAGCAGTCAGCTCCAGTCTCCAAAGTCTCTAGATGT TCTGCCCTCCAAAGTCTATTCTGGTGGAGCAGTCCCTCCGACCCACATCTCTTA TAATTCTTGCCCTGTCCAAGATCAGCATGCCATCATCATTCCAGATACTCCAG CCCTCCTGTGAGTGTCACTATCCGAAGTGAACACTGATGAGGAAGAGGACAACAAATA CAAGCCCAGTAGCTCTGGACTGAAGCCAAGGTCTAATGTCACTGTTATGTCAGTCAA TGATTCTCCAGACTCTGACTCTTGTGAGCAGCCCTATTCCACTGATACCCGTAGTGC TCTCCGAGGCAATAGTGGATCCGTTGGAGGGGCTGGCAGAGTTGTGGCAGATGGCAC TGGCACCCGCACTATCATTGCGCTTCACTGAAAACCTCAGCTGGTACTGCACTGTAGC AACCCAGGCCCTCAGGTCTCCGTGAGCAATAAGACTAAGCCAGTCGCTTCAGTGAGGGCA GTCATCTGGATGCTGTATCACCCCCACAGGTATCGAGCTAACGCCGGGGGACCAAGTGC AGCACAACCACCAATCTAGCCAGAACCCAGCAGTCAGTCAGTCAGCTCCACAGGA GAGAAGCAGCAACCCAGCCCCCGCAGGCAGCAGGGCTTGTGGCCCTCTCCCAAGC CCCCTACACCTCCAGCATGGCAGCCGCTACACTCGACAGGGCACCCACACCTGCC GGCCCTGCTACCTGCCAGGCCAGGCTCATCTGATACGTATGTCGCCCCGACTCTGC TGCTGCACTGGCTCAACCGAGCTCCATTGCTCATCTTCTCCCCACAGGGTCTCAAG GCATGCTGAGCCTATACCACTCACCTAGCACTTGGTGCACCAGGCTCTGCACTG TGGCCCAGCCTCTCACTCTGCCAGCGTGGCCCTGCTCAGTACCAACACCAAGTTGC CACCAATCCTACATTGGTCTCCGAGGCTAACAAATTACACTGGATACCCGCTGAG TCCTACCAAGATCAGCCAGTATTCTACTTATAGTTGGTGAGCATGAGGGAGGAGGAATC ATGGCTACCTCTCTGGCCCTGCGTTCTTAATTGGGCTATGGAGAGATCCTCTTAA CCCTCTGAAATTCTTAGCCAGCAACTGTTCTGCAGGGCCCACTGAAGCAGAAGGTT TTCTCTGGGGGAAACCTGCTCAGTGGTACTGCATTGTTGAGTCTCCCAAAGTTGC CCTATTTAAATTCTTGTGACAGTAATTGGTACTGGAAAGAGTTCAAGTGAAGCCCCCT CCCATCTCTGCACTTACCAAGGAAGAGAGATTGTTCTGAAGTACCCCTGAAAAATAT TTTGTCTCTGACTTGATTCTATAAATGCTTTAAAAACAAGTGAAGCCCCCTTTAT</p>

		TTCACTTGTGTTATTGTGATTGCTGGTCAGGAAAAATGCTGATAGAAGGAGTTGAAATC TGATGACAAAAAAAAGAAAAATTACTTTTGTGTTATAAACCTCAGACTGCCTATT ATTTAAAAGCGGCCTAACACAATCTCCCTTGTGTTATTGGACATTAAACTACAGAGT TTCAGTTTGTGTTAATGTCATATTACTTAATGGGCAATTGTTATTGCAAACACTG GTTACGTATTACTCTGTGTTACTATTGAGATTCTCAATTGCTCCTGTGTTGTTATAA AGTAGTGTGTTAAAAGGCAGCCTACCATTTGCTGGTACTTAATGTGAGAGAATCCATATC TGCCTGAAAACACCAAGTATTCTTTAAATGAAGCACCCTGAATTCTTTAAATTAT TTTTAAAAGTCTTCTCTGATTCACTAAATTGAAAGCTTAACTTCAAAAGTAGCTTCCCT TGTATGCCAGCAGCAAATTGAATGCTCTCTTAAAGACTTATAATAAGTGCATGTAG GAATTGCAAAAATATTAAAATTACTGAATTAAAATATTAGAAGTTTG TAATGGTGGTGTAAATTACATAATTAAATATGTACATATTGATTAGAAAATAT AACAAAGCAATTCTGCTAACCCAAAATGTTATTGTAATCAAATGTGAGTGTGATTAC ACTTGAATTGTGACTTAGTGTGTTAGCTGATCCTCCAGTGTATCCCAGGATGGATTGA TGTCTCCATTGTATTAAACCAAAATGAACTGATACTTGTGGAATGTATGTGAACTAAT TGCAATTATATTAGAGCATATTACTGTAGTGTGCTGAATGAGCAGGGGCTTGCCTGCAAGG AGAGGAGACCCCTGGAATTGTTGCACAGGTGTGCTGGTGGAGGAGTTTCAGTGTG GTCTCCCTCCCTTCTCCCTTATTGTAAGTGCCTTATATGATAATGTAGT GGTTAATAGAGTTACAGTGTGAGCTGCCTAGGATGGACCAGCAAGCCCCGTGGACCC AAGTTGTTCACCGGGATTATCAGAACAGGATTAGTAGCTGTATTGTGTAATGCATTGTT CTCAGTTCCCTGCCAACATTGAAAATAAAACAGCAGCTTCTCCTTACCAACACC TCTACCCCTTCCATTGGATTCTGGCTGAGTTCTCACAGAACGATTCCCCATGTG GCTCTCACTGTGCGTTGCTACCTTGTGAGAATTCAAGAACGAGGTGAGGAGGA GTCAAGCCAATATTAAATATGCAATTAAAGTATGTGCAATCACTTAAAGTGAAT TTTTTCTTCCCATGTGGCAGTCCCTGCACATAGTTGACATTCTAGTAAAA TATTGCTTGTGAAAAAAACATGTTAACAGATGTGTTATACAAAGAGCCTGTTGAT TGCTTACCATGTCCCCATACTATGAGGAGAAGTGTGCTGGTGCCTGGTACAAGGAAC TCACAGAAAGGTTCTAGCTGGTGAAGAATATAGAGAACCAAGCCTGTTGAGTC ATTGAGGCTTTGAGGTTCTTTAAACAGCTGTATAGTCTGGGGCCCTCAAGCTG TGAAATTGTCTTGACTCTCAGCTGCTGATGGATCTGGTCAAGTAGAGGTACTGG GATGGGGACATTCCCTGCCATAAAGGATTGGGAAAGAACGATTAATCCTAAAATACAGG TGTGTTCCATCCGAATTGAAAATGATATATTGAGATATAATTAGGACTGGTTCTGTG TAGATAGAGATGGTGTCAAGGAGGTGCAGGATGGAGATGGGAGATTTCATGGACCTGGT CAGCCAGCTGTACCAGGTTGAACACCGAGGAGCTGCAAAGTATTGGAGTTCTCA TTGTAAGGAGTAAGGGCTCCAAGATGGGCAGGTAGTCCGTACAGCCTACCAAGGAACAT GTTGTGTTCTTATTTAAAATCATTATATTGAGTTGTGTTCTGAGCACTATATTG GTCAAGATAGCCAAGCAGTTGTATAATTCTGCACTAGTGTCACTACAGTTCTGGTC AACATGTGTGATCTTGTGCTCTTTGCCAACGACATTCTGATTTCTGTTGGAAC ACAGGTCTAGTTCTAAAGGACAAATTGTTCTGCTGCAAGCAGCATTCTGATTTCTGTTG GATTGTTGTTGTAAGGAAATGAGATGCAGGAAAGAAAACCAAATCCATTCCCTGCAC CCCAGTCCAATAAGCAGATAACCAACTTAAGATAGGAGTCTAAACTCCACAGAAAAGGATAA TACCAAGAGCTGTATTGTTACCTAGTCACCTGCCTAGCAGTGTGTTGCTTAAACT AGAGATTTCAGTCTAGCTGCAAACCTGGCATTCCGATTTCAGCATAAAATCCA CCTGTGCTGCTGAATGTGTTATGTGCTACTGTGCTTAAAGGCTTAAAGTCCCTGGG TTAGCCCTGTTGGCCCTGACAGGAAGGGAGGAAGCCTGGTGAATTAGTGAAGCAGCTGG CTGGGTACAGTGACCTGACCTCAAACCAAGCTTAAGGCTTAAAGTCCCTCTCAGAACTT
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			GGCATTCCAACCTCTCCCTTCCGGGTGAGAGAAGAAGCGGAGAAGGGTCAGTGTAGC CACTCTGGCTCATGGGACACTTGGTCACTCCAGAGTTTAATAGCTCCCAGGAGGTG ATATTATTTCACTGCTCAGCTGAAATACCAACCCAGGAATAAGAACCTCCATTCAAAC AGTTCTGCCATTCTGAGCCTGCTTGTGATTGCTCATCCATTGCTCCACTAGAGGG GCTAAGCTTGACTGCCCTAGCCAGGCAAGCACAGTAATGTGTGTTGTTAGCATTAT TATGCAAAAATTCACTAGTTGAGATGGTTGTTAGGATAGGAAATGAAATTGCCTCTC AGTGACAGGAGTGGCCCGAGCCTGCTCCTATTGATTTTTTTAACTGATAG ATGGTGAGCATGTCTACATGGTGTGCTAAACTTATATAATGTGTGGTTCAA TTCAGCTGAAAAATAATCTCACTACATGTAGCAGTACATTATGTACATTATGTAA TGTTAGTATTCGCTTGAAATCCTGATATTGCAATGGAATTCCCTACTTTATTAAATGT ATTTGATATGCTAGTTATTGTCGATTAACTTTTGCTTCTCCCTTTTTGG TTGTCGCTTCTTACAACAAGCCTCTAGAAACAGATAGTTCTGAGAATTACTGAGC TATGTTGTAATGCAGATGACTTAGGGAGTATGAAAATAATCATTTAACAAAAGAAA TAGATATTAAAATTAACTAACTATGGGAAAGGGTCCATTGTGTAAAACATAGTT ATCTTGGATTCAATGTTGCTTGGTTTACAAGTAGCTGTATTTCACTGATTTC TACATAATATGGAAAATGTAGAGCAATTGCAATGCATCAATAAAATGGTAAATTCT G
S000023	F31	158	GGAGCCGTACCCCCGGGGGGGACCCAGCGCAGGCACTCCCGCGGGCGCCCGGAGG GAGGGAGCGAGCGGGCGGGCGGGCAAGCCAGACAGCTGGCCGGAGCAGCCGGCGGCC CGAGGGGCCAGCGAGATTGTAACCATGGCTGTGAGTACAAGCTCAGCAGCTCAAAG GAGAAGCCCTTCATCAGATGCAAGCGTTATATGCCAGCATTTCCATTGAGGTGCGGC ATTATTATCCCAGTGGATTGAAAGCCAAGCATGGACTCAGTAGATCTGATAATCCAC AGGAGAACATTAAGGCCACCCAGCTCCTGGAGGGCCTGGTCAGGAGCTGCAGAAGAAGG CAGAGCACCAGGTGGGGAAGATGGTTTACTGAAGATCAAGCTGGGCACTATGCCA CACAGCTCCAGAACACGTATGACCGCTGCCCCATGGAGCTGGCCGCTGCATCCGCCATA TATTGACAATGAACAGAGGTTGGTCGAGAAGCCAACAATGGTAGCTCTCCAGCTGGAA GCCTGCTGATGCCATGTCAGGACACAGAGAATGAGTTAAAAGCTGCAGCAGACTCAGGAGTACT TCATCATCCAGTACCAAGGAGAGCCTGAGGATCCAAGCTCAGTTGGCCGCTGGCCAGC TGAGCCCCAGGAGCGCTGAGCCGGAGACGGCCCTCCAGCAGAACAGCTGGAGCTCTGG AGGCCTGGTTGAGCGTGAAGGACAGACACTGCAGCAGTACCGCGTGGAGCTGCCAGA AGCACCAAGAACCTGCAAGCTGCTGCCAGCAGACCATCATCCTGGATGACGAGC TGATCCAGTGGAAAGCGGGCAGCAGCTGGCCGGAAACGGCGGGCCCCAGGGCAGCC TGGAACGTGCTACAGTCCTGGTGAGAAGTTGGCGAGATCATCTGGCAGAACCGGCAGC AGATCCGCAGGGCTGAGCACCTCTGCCAGCAGCTGCCATCCCCGGCCAGTGGAGGAGA TGCTGGCCAGGGTCAACGCCACCATCACGGACATTATCTCAGCCCTGGTACCAGCACGT TCATCATGGAGAACAGCAGCCCTCTCAGGTCTGAAGACCCAGACCAAGTTGCAGCCACTG TGCGCCTGCTGGTGGGGGGAGCTGAACGTGCACATGAACCCCCCCCAGGTGAAGGCCA CCATCATCAGTGAACAGCAGGCCAGTCTGCTCAAGAACAGAGAACACCCGCAATGATT ACAGTGGCGAGATCTTGAACAACACTGCTGCGTCAAGGAGTACCAAGGCCACAGGCACCC TTAGTGCCTCAGGAATATGCTCTGAACAGAATTAAAGAGGTCAAGACCGTCTGGGG CAGAGTCGGTACAGAAGAAAAATTACAATCTGTTGAATCCCAAGTTCAGTGTGG GAAATGAGCTGGTTTCAAGTCAAGACCCCTGCCAGTGGTGTGATGTTCTGAGAGCCTG GCAGCCAGGACAACAATGCGACGCCACTGTTCTCTGGGACAATGCTTGGAGGCC GCAGGGTGCCTTGCCTGACAAAGTGCAGAGCAACGGGGCTGACCAAGGAGAACCTCGT ACATGAAATTCAAGGCCAGTGCAGAGCAACCCGGGGCTGACCAAGGAGAACCTCGT TCCTGGCCAGAAACTGTTCAACACAGCAGCACCCAGGAGACTACAGTGGCTGT CTGTGCTCTGGCCCAGTTCAACAGGGAGAATTACCAAGGAGCGAATTACACTTCTGGC AATGGTTGACGGTGTATGGAAGTGTAAAAAAACATCTCAAGCCTATTGGAATGATG

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			GGGCCATTTGGGTTGTAAACAAAGCAACAGGCCATGACCTACTGATTAACAAGCCAG ATGGGACCTCCTCCTGAGATTCACTGACTCAGAAATTGGCGCATCACCATGCTTGGA AGTTGATTCTCAGGAAAGAATGTTTGAATCTGATGCCTTACCAACCAGAGACTCT CCATCAGGTCCCTAGCCGACCGCTGGAGACTGAATTACCTATCTACGTCTTCTG ATCGGCCAAAGATGAAGTATACTCCAAACTACACACCCAGTCCCTGCGAGTCGCTA CTGCTAAAGCTGTGATGGATACGTGAAGCCACAGATCAAGCAAGTGGTCCCTGAGTTG TGAAACGCATCTGCAGATGCCGGGGCGGCAGCGCCACGTACATGGACCCAGGCCCCCTCCC CAGCTGTGTGCCCCAGGCTCACTATAACATGTACCCACAGAACCTGACTCAGTCCTG ACACCGATGGGACTTCGATCTGGAGGACACAATGGACGTAGCGCGCGTGGAGGAGC TCCTGGGCCGCAATGGACAGTCAGTGGATCCGCACGCACAATCGTGAACCCCGCACC TCTCCATCTCAGCTCTTCATCTTCAACCAGAGGAATCACTCTGTGGATGTTTAATT CATGAATCGCTCTCTTTGAAACAATACTCATAATGTGAAGTGTAAATACTAGTTGTGA CCTTAGTGTCTGTGCATGGTGGCACCGCGAAGGGAGTGCAGTATGTGTTGTGT GTGTGTGTGTGTGTGTGCGTTGGTCACGTTATGGTGTGTTCTCCCTCACTGT CTGAGAGTTAGTTGTAGCAGA
S000031	F32	159	CCGAATGTGACCGCCTCCGCTCCCTCACCGCCGGGGAGGAGGAGCGGGCGAGAAC TGCCGCCGAACGACAGGACGTTGGGCGGCCCTGGCTCCCTCAGGTTAACGATGTTAA GCTGCATCAATGGAGCACATACAGGGAGCTTGAAGACGATCAGCAATGGTTGGATT AAAGATGCCGTGTTGATGGCTCCAGCTGCATCTCCTACAATAGTCAGCAGTTGGC TATCAGGCCGGGCATCAGATGATGGCAAACACTCACAGATCCTCTAAAGACAAGCAACACT ATCCGTTTCTGCCAACAGCAAAGAACAGTGGTCAATGTGCGAACATGGAAATGAGC TTGCATGACTGCCATTGAAAGCACTCAAGGTGAGGGGCCTGCAACCAGAGTGTGTGCA GTGTCAGACTCTCCACGAACACAAAGGTAAGGACAGCTTAGATTGAAATACTGAT GCTGCGTCTTGATTGGAGAACAGTCAAGTAGATTCCCTGGATCATGTTCCCTCACA ACACACAACCTTGCCTGGAAAGACGTTCTGAAGCTGCCTCTGTGACATCTGTCA TTCCGTCAATGGATTGATGTCAGACTGTGGCTACAAATTCTCATGAGCACTGTAGC ACCAAAGTACCTACTATGTGTGGACTGGAGTAACATCAGACAACCTTATTGTTCCA AATTCCACTATTGGTATAGTGGAGTCCCAGCACTACCTCTTGACTATGCGTGT CGAGAGTCTGTTCCAGGATGCCTGTTAGTTCTCAGCACAGATATTCTACACCTCAGCC TTCACCTTAACACCTCCAGTCCCTCATCTGAAGGTTCCCTCCAGAGGCAGAGGTCG ACATCCACACCTAATGTCCACATGGTCAGCACACGCTGCTGTGGACAGCAGGATGATT GAGGATGCAATTGCAAGTCACAGCGAATCAGCTCACCTTCAGCCCTGTCCAGTAGCCCC AACATCTGAGCCCAACAGGCTGGTCACAGCCAAAACCCCCGTGCCAGCACAAGAGAG CGGGCACCAAGTATCTGGACCCAGGAGAAAAACAAAATTAGGCCCTGGACAGAGAGAT TCAAGCTATTGGAAATAGAAGCCAGTGAAGTGTGCTGCCTCGGATTGGTCA GGCTTTGGAACTGTTATAAGGGTAAATGGCACGGAGATGTTGAGTAAAGATCCTA AAGGGTGTGACCCAACCCAGAGCAATTCCAGGCCCTCAGGAATGAGGTGGCTGTTCTG CGCAAAACACGGCATGTGAACATTCTGTTTCACTGGGTCATGACAAAGGACAACCTG GCAATTGTGACCCAGTGGTGCAGGGCAGCAGCCTCTACAAACACCTGCATGTCCAGGAG ACCAAGTTCAGATGTTCCAGCTAATTGACATTGCCGGCAGACGGCTCAGGGAAATGGAC TATTGATGCAAAGAACATCATCCATAGAGACATGAAATCAAACAATATATTCTCCAT GAAGGCTTAACAGTGAAGGAGATTGGTTGGCAACAGTAAAGTCACGCTGGAGT GGTCTCAGCAGGTTGAACAACCTACTGGCTCTGTCTGGATGGCCCCAGAGGTGATC CGAATGCAGGATAACAACCCATTCACTGTTCCAGTGGATGTCTACTCCATGGCATCGTA TTGTATGAACTGATGACGGGGAGCTCCATTCTCACATCAACAACCGAGATCAGATC ATCTCATGGGGCCAGGATATGCCCTCCAGATCTTAGTAAGCTATATAAGAAACTGC CCCAAAGCAATGAAGAGGCTGGTAGCTGACTGTGTGAAGAAAAGTAAAGGAAGAGAGGCCT CTTTTCCCCAGATCCTGCTCCATTGAGCTGCTCCAACACTCTCACCGAAGATCAAC CGGAGCGCTCCGAGCCATCCTGCACTGGCAGCCCACACTGAGGATATCAATGCTTGC

			ACGCTGACCACGTCCCCGAGGCTGCCGTCTTAGTTGACTTGACCTGTCTCAGGC TGCCAGGGGAGGAGGAAGCCAGCAGGCACCACTTCTGCTCCCTTCTCCAGAGGA GAACACATGTTTCAGAGAAGCTGCTAAGGACCTCTAGACTGCTCACAGGGCTAA CTTCATGTTGCCCTCTTCTATCCCTGGGCCCTGGAGAAGGAAGCCATTGAGTG CTGGTGTCTGCTCCCTCCCACATCCCCATGCTCAAGGCCAGCCTCTGTAGATG CGCAAGTGGATGTTGAGTAGTACAAAAGCAGGGCCAGCCCCAGCTGTTGGCTACA TGAGTATTTAGAGGAAGTAAGGTAGCAGGCAGTCCAGCCCTGATGTTGGAGACACATGGGA TTTGGAAATCAGCTTCTGGAGGAATGCATGTCACAGGGGGACTTCTCAGAGAGTGG TGCAGCGCCAGACATTTGCACATAAGGCACCAAACAGCCCAGGACTGCCAGACTCTGG CCGCCCCAAGGGAGCCTGCTTGGTACTATGGAACCTTCTAGGGACACGTCCCTTT CACAGCTCTAAGGTGTCCAGTGCACTGGGATGGTTCCAGGCAAGGCACCTGGCAAT CCGCATCTCAGCCCTCTCAGGAGCAGTCTTCCATCATGCTGAATTGTCTCAGGAGC TGCCCCTATGGGCGGGCCAGGGCCAGCCTGTTCTAACAAACAAACAAACA GCCTGTTCTAGTCACATCATGTTATAAGGAAGCCAGGAATACAGGTTCTG ATGATTTGGTTTTAATTGTTTATTGACCTGACAAACACAGTTATCTGATGGTC CCTCAATTATGTTATTAAATAAAATAATTAAATT
S000039	F33	160	TCCAGTTGCTCTGGAGAACACTGGACAGCTGAATAATGCAGTATCTAAATATAAA GAGGACTGCAATGCCATGGCTTCTGTGCTAAATGAGGAGCTCCAAGAACAGACTGAGGTG AACCTGGAGGGCCCTGAGCCAGGGTGGAGTGATCTCTATCTGCGACAGGGAGCCC CTCCGGCTGGCAGTGGAGAGTACACAGCAGAGGAACCTGTGCATCAGGCTGCACAGGCA TGCCGTATCTCCTCTTGTACAAACCTCTTGCCTGTATGACGAGAACACCAAGCTC TGGTATGCTCCAAATCGCACCATACCGTTGATGACAAGATGTCCTCCGGCTCCACTAC CGGATGAGGTTCTATTCAACCAATTGGCATGGAACCAACGACAATGAGCAGTCAGTGTGG CGTCATTCTCCAAAGAACAGAGAAAAATGGCTACGAGAAAAAAAGATCCAGATGCAACC CCTCCTTGATGCCAGCTCACTGGAGTATCTGTTGCTCAGGGACAGTATGATTGGTG AAATGCTGGCTCTATTGAGACCCCAAGACCGAGCAGGATGGACATGATATTGAGAAC GAGTGTCTAGGGATGGCTGCTGGCATCTCACACTATGCCATGATGAAGAACATGAG TTGCCAGAACTGCCAAGGACATCAGTACAAGCGATATATTCCAGAAACATTGAATAAG TCCATCAGACAGAGAACCTCTCACCAGGATCGGATAAAATGTTCAAGGATTTC CTAAAGGAATTAAACAACAAAGACCATTGTGACAGCAGCGTGTCCACGCATGACCTGAAG GTGAAATACTGGCTACCTTGAAACCTTGACAAAACATTACGGTGTGAAATATTGAG ACTTCATGTTACTGATTCATCAGAAAATGAGATGAATTGGTTCTTCATGACGGT GGAAACGTTCTACTACCGAAGTGATGGTACTGGGATCTGGAAATCCAGTGGAGGCAT AAACCAAATGGTTCTGTTGAAAGGAAAAAAACTGAAGCGAAAAAAACTGGAA AATAAGACAAGAAGGATGAGGAGAAAAACAAGATCCGGAAAGAGTGGAAACAATTTC TTCTCCCTGAAATCACTCACATTGTAATAAAGGAGTCTGTTGTCAGCATTAAACAGCAG GACAACAAGAAATGAACTGAAGCTCTTCCACGGAGGCTTGCTCTTG CTGGTAGATGGCTACTTCCGGCTCACAGCAGATGCCATATTACCTCTGCACCGACGTG GCCCGGGCTGATGTCACACATACAGAATGGCTGTCATGGTCCAATCTGTACAGAA TACGCCATCAATAAATTGCGCAAGAAGGAAGCGAGGAGGGATGTACGTGCTGAGGTGG AGCTGCACCGACTTGACAAACATCCTCATGACCGTCACCTGCTTGGAGAAGTCTGAGCAG GTGAGGGTGGCCAGAAGCAGTTCAAGAACTTCAAGATCGAGGTGCAAGAAGGGCGTAC AGTCTGCACGGTTCGGACCGCAGCTCCCCAGCTGGAGACCTCATGAGCCACCTCAAG AAGCAGATCCTGCGCACGGATAACATCAGCTTCTGCTAAACGCTGCTGCCAGCCCAAG CCCCGAGAAATCTCCAACCTGCTGGGCTACTAAGAAAGCCAGGAGTGGCAGCCGTC TACCCCATGAGCCAGCTGAGTTGATGGATCCTCAAGAAGGATCTGGTGCAGGGCGAG CACCTGGGAGAGGACGGAGAACACACATCTATTCTGGACCCCTGATGGATTACAAGGAT GACGAAGGAACCTCTGAAGAGAAGAAGATAAAAGTGTACCTCAAAGTCTTAGACCCAGC CACAGGGATATTCCCTGGCCTTCTCGAGGCAGCCAGCATGATGAGACAGGTCTCCAC

			AAACACATCGTGTACCTCATGGCGTCTGTGTCGCCGACGTGGAGAATATCATGGTGGAA GAGTTGTGGAAGGGGGCCTCTGGATCTTCTCATGCACCGGAAAAGTGTGTCCTTAC ACACCATGGAAATTCAAAGTGCACAGCTGCCAGTGCCCTGAGCTACTGGAGGAT AAAGACCTGGCATGGAAATGTGTACTAAAAACCTCCCTGGCCGTGAGGAAATC GACAGTGAGTGTGGCCATTCAAGCTCAGTGACCCGGCATCCCCATTACGGTCTG TCTAGGCAAGAATGCATTGAACGAATCCCAGTGGATTGCTCCTGAGTGTGTTGAGGACTCC AAGAACCTGAGTGTGGCTGCTGACAAGTGGAGCTTGGAAACCACGCTCTGGGAAATCTGC TACAATGGCGAGATCCCTGAAAGACAAGACGCTGATTGAGAAAGAGAGATTCTATGAA AGCCGGTGCAGGCCAGTGACACCATCATGTAAGGAGCTGGCTGACCTCATGACCCGCTGC ATGAACATGACCCCAATCAGAGGCCCTTCTCCGAGCCATCATGAGAGACATTAATAAG CTTGAAGAGCAGAACATCCAGATATTGTTCCAGAAAAAAACAGCCAACCTGAAGTGGAC CCCACACATTTGAGAAGCGCTCCTAAAGAGGATCCGTGACTTGGGAGAGGGCCTTT GGGAAGGTTGAGCTGAGGTATGACCCGAAGACAATACAGGGAGCAGTGGCTGTT AAATCTGAAAGCCTGAGAGTGGAGGTAACCACATAGCTGATCTGAAAAAGGAAATCGAG ATCTTAAGGAACCTCATGAGAACATTGTGAAGTACAAAGGAATCTGCACAGAAC GGAGGAATGGTATTAGCTCATGGAATTCTGCCTCGGGAGCCTTAAGGAATAT CTTCCAAAGAATAAGAACAAAATAACCTCAAACAGCAGCTAAATATGCCGTTAGATT TGTAAGGGATGGACTATTGGTTCTCGGCAATACGTTCACCGGGACTTGGCAGCAAGA AATGTCCTGTTGAGAGTGAACACCAAGTGAAGACTTCGGTTAACAAAGCA ATTGAAACCGATAAGGAGTATTACACCGTCAAGGATGACGGGGACAGCCCTGTGTTTGG TATGCTCCAGAATGTTAATGCAATCTAAATTATATTGCCCTGACGCTGGTCTTT GGAGTCACTCTGCATGAGCTGCTGACTTACTGTGATTAGCTAGTCCCCTGGCTT TTCTGAAAATGATAGGCCAACCCATGGCCAGATGACAGTCACAAGACTTGTGAATACG TTAAAAGAAGGAAACGCCGTGCCGTGCCACCTAACTGTCCAGATGAGGTTATCAGCTT ATGAGAAAATGCTGGAAATTCAACCATCCAATCGGACAAGCTTCAAGAACCTTATTGAA GGATTGAAAGCACTTTAAAATAAGAACATGAATAACATTAAATTCCACAGATTATCA A
S000040	F34	161	CTGCAGCTCTAGGACCCGGTTCTTACTGATTAAAAACAAAACAAAAAAATAAA AAAGTTGTGCCTGAAATGAATCTGTTTTTTTTATAAGTAGCCGCCTGGTTACTGTG CCTGAAAATACAGACATTGACCCCTGGTGTAGCTCTGTCACCTTATATCACGGAA TGGATGGGCTGATTCTGGCCCTCTCTGAATTGGCCATATACAGGGTCCCTGGCCA GTGGACTGAAGGCTTGTCAAGATGACAAGGGTCAGCTCAGGGATGTGGGGAGGGCG GTTTATCTCCCCCTGTGCTTGTAGGTTGATCTCTGGTAAGAGGGCGTTATCT TTGTAACACGAAACATTGTCTCCAGTTCTGTTAATGGCAGAACATGGAAAG CGAATAAAGTTACTGATTTTGAGACACTAGCACCTAGCGCTTCTATTGAAACGT CCCGTGTGGAGGGCGGGCTGGGTGCGGCTGCCGATGACTCGTGGTTGGAGGGCCA CGTGGCCGGGGCGGGACTCAGGCCCTGGCAGCCACTGATTACGTAGCGGGGGCG GGAAGTGCCGCTCTGGGGCTGTTCATGGCGTCCGGGCTCCAAACATTTC CCGGTCTGTGGCCTAAATCTGTCAAAGCAGAGGCAGTGGAGCAGGTGGTGGAAAAGCGCA TGTGAAATGACTGAGTACAAACTGGTGGTGGAGCAGGTGGTGGAAAAGCGCA CTGACAATCCAGCTAACCAAGAACACTTGTAGATGAATATGATCCCACCATAGAGGAT TCTTACAGAAAACAAGTGGTATAGATGGTGAACCTGTTGGACATACTGGATACA GCTGGACAAGAAGAGTACAGTGCCTGAGAGACCAATACATGAGGACAGGCCAGGCTTC CTCTGTGATTGCCATCAATAAGCAAGTCATTGCGGATATTAACCTCTACAGGGAG CAGATTAAGCGAGTAAAGACTCGGATGATGTAACCTATGGTCTAGGGAAACAAGTGT GATTGCCAACAGGACAGTGTGATACAAAACAAGCCCACGAACTGGCCAAGAGTACGGG ATTCCATTGAAACCTCAGCCAAGACCAGACAGGGTGTGAAGATGCTTTACACA CTGGTAAGAGAAAACGCCAGTACCGAATGAAAAAACTCAACAGCAGTGTGATGGACT CAGGGTTGTATGGATTGCCATGTGTTGATGTAACAGATACTTTAAAGTTGTCA

			GAAAAGAGCCACTTCAAGCTGCACTGACACCCGGTCCTGACTTCTGGAGGAGAAGTA TTCCCTGTTGCTGTCTTCAGTCTCACAGAGAACGCTCTGCTACTTCCCCAGCTCTCAGTAG TTAGTACAATAATCTCTATTGAGAAGTCTCAGAATAACTACCTCTCACTTGGCTGT CTGACCAGAGAACATGCACCTCTGTTACTCCCTGTTATTTCTGCCCTGGGTTCTCCAC AGCACAAACACACCTCAACACACCTCTGCCACCCCAGGTTTCATCTGAAAAGCAGTTC ATGCTGAAACAGAGAACCAAACCGCAAACGTGAAATTCTATTGAAAACAGTGTCTGAG CTCTAAAGTAGCAACTGCTGGTGATTTTTTTCTTTTACTGTTGAACCTAGAACTAT GCCTAATTTGGAGAAATGTCTAAATTACTGTTGCAAGAATATAGTTATTATTGC TGTGTTGGTTGTTATAATGTTATCGGCTCTATTCTCTAAACTGGCATCTGCTCTAGATT CTAAATACAAAATGAATACTGAATTGGAGTCTATCCTAGTCTTCAAACTTGTACGT AATTAATCCAACCTTCACAGTGAAGTGCCTTTCTAGAAGTGGTTGTAGACTCCT TTATAATATTCAGTGGAAATAGATGTCCTAAACCTTATGCATGAAATGAATGTCTGA GATACGTCTGTGACTTATCACCATTGAAGGAAAGCTATATCTATTGAGAGCAGATGCC ATTTGTACATGTATGAAATTGGTTTCCAGAGGCCGTGTTGGGCTTCCCAGGAGAA AGATGAAACTGAAAGCATATGAATAATTCACTTAAATTTACCTAATCTCCACTTT TTTCATAGGTTACTACCTATACAATGTATGTAATTGTTCCCTAGCTTACTGATAAAC CTAATATTCAATGAACCTCCATTGTATTCAAATTGTGTATACCAGAAAGCTCTACAT TTGAGATGTTCAAATATTGAAAACCTGGTGATTGTTATTAATAGCTGTGATCAGT GATTTCAAACCTCAAATATAGTATATTAAACAAATT
S000046	F35	162	CGGGGGATCTGGCTGTTGCTGGATCTGTAGTGGCGGGCGGGCGGGCGGGCGGG GGAGGCAGCAGGCGCGGGAGCGGGCGCAGGAGCAGGCGGGCGGTGGCGGGCGGGTTA GACATGAACGCCGCCTCGCGCCGGCGGTGACGGAGAGGCCCTCTCGCGCGGGCGGG TTGTGTGATTTGCTAAATGCATACCAACAGCGAATGGCTGCCTAGGGACGGACAA AGAGCTGAGTGTATGGATTCTAGTGCATGTTCACCTCCTGTGAGCAGTGGGAA AAATGGACCAACTTCTGGCAAGTGGACATTTACTGGCTCAAATGAGAAGACAGAAG TAGCTCAGGGCTCTGGGGATGGAGGACATCCAAGCCGTCAGGAACATGGAGATGG GACTCCCTATGACCACATGACCAGCAGGGACCTGGTACATGACAATCTCTCCACC TTTGTCAATTCAGAATACAAAGTAAACAGAAAGGGGCTCATACTCATCTTATGGAG AGAATCAAACCTACAGGTTGCCACCAGCAGAGTCTCTGGAGGTGACATGGATATGG CAACCCAGGAACCCCTTCGCCCACCAAACCTGGTCCCAGTACTATCAGTATTCTAGCAA TAATCCCCGAAGGGAGGCTTTCACAGTAGTGCATGGAGGTACAGACAAAGAAAGTTCG AAAAGTTCTCCAGGTTGCCATCTCAGTCTATGCTCCATCAGCAAGCACTGCCACTA CAATAGGGACTGCCAGGCTATCCTCTCCAAACCAGCAACCAGCAGTCCAGTGGGATGAA CTTCTCATGCAAGATGCCATCACAGCAGTGCACCTGGAGCTCTCCAGTGGGATGAA TCAGCCTGGCTATGCAGGAATGTTGGCAACTCTCTCATATTCCACAGTCCAGCAGCTA CTGAGCCTGCATCCACATGAACGTTGAGCTATCCATCACACTCCTCAGCAGACATCAA TTCCAGTCTCTCCGATGTCACCTCCATCGTAGTGGTACAAACCAATTACAGCACCTC TTCTGTACGCCCTGCCAACGGGACAGACAGTATAATGGCAAATAGAGGAAGCGGGGG AGCCGGCAGCTCCAGACTGGAGATGCTCTGGGGAAAGCAGTCTCGATCTATTCTCC AGATCACACTAACACAGCTTTCATCAAACCCCTCAACTCCTGTTGGCTCTCCATC TCTCTCAGCAGGACAGCTGTTGGCTAGAAATGGAGGACAGGCCATCGTCTCTAA TTATGAAGGACCTTACACTCTTGCAAAGCCGAATTGAAGATGTTAGAAAGACTGGA TGATGCTATTCTGTTCTCCGGAACCATGCAGTGGGCCATCCACAGCTATGCTGGTGG TCATGGGACATGCATGGAATCTGGACCTCTCATATGGAGCCATGGTGGCTGG CTCAGGGTATGGAACCGGCCCTCTTCAGCCAACAGACATTCACTCATGGTGGGGACCCA TCGTGAAGATGGCGTGGCCCTGAGAGGGCAGCCATTCTCTGCCAAACCAGGTTCCGGT TCCACAGCTCTGTCCAGTCTGCAGTCTCCCCCTGACCTGAACCCACCCAGGACCCCTA CAGAGGCATGCCACCAGGACTACAGGGGAGCAGAGTGTCTCTGGCAGCTGAGATCAA ATCCGATGACGAGGGTGTAGAGAACCTGCAAGACACGAAATCTCGGAGGACAAGAAATT

			AGATGACGACAAGAAGGATATCAAATCAATTACTAGCAATAATGACGATGAGGACCTGAC ACCAAGAGCAGAAGGCAGAGCGTGAGAAGGGCGGAGGATGCCAACATGCCGAGAGCG TCTGGGGTCCGTGACATCAACGAGGCTTCAAAGAGCTCGCCGATGGTCAGCTCCA CCTCAAGAGTGACAAGCCCCAGACCAAGCTCCTGATCCTCACCAGGGCGTGGCGTCAT CCTCAGTCTGGAGCAGCAAGTCCGAGAAAGGAATCTGAATCCGAAAGCTGCGTGTCTGAA AAGAAGGGAGGAAGAGAAGGTCTCGAGCCTCCCCCTCTCCTTGGCGGCCACAC CCCTGGAATGGGAGACGCATCGAATCACATGGGACAGATGTAAGGGTCCAAGTTGCCA CATTGCTTCACTAAACAGAGACCACTCCTAACAGCTGATTATCTAAACCCACAT AAACACTTCTCCTAACCCCCATTTGTAATATAAGACAAGTCTGAGTAGTTATGAATC GCAGACGCAAGAGGTTTCAGCATTCCAATTATCAAAAAACAGAAAAACAAAAAAAGAA AGAAAAAAAGTGCACCTTGAGGGACGACTTCTTAAACATATCATTCAAATGTGCAAAGC AGTATGTACAGGCTGAGACACAGCCAGAGACTGAACGGC
S000050	F36	163	AAAAAAAAAGAAAAAAAAGGCACAAAAAGTGGAAACTTTCCCTGTCATTCCATCAAG TCCTGAAAATCAAAATGGATTAGAGAAAAATTATCCGACTCCTCGGACCAGCAGGACA GGACATGGAGGAGTGAATCAGCTGGGGGGTTTTGTGAATGGACGGCCACTCCGGAT GTAGTCGCCAGAGGATAGTGGAACTTGCTCATCAAGGTGTCAAGGCCTGCGACATCTCC AGGCAGCTTGGGTAGCCATGGTTGTCAAGCAAATTCTGGCAGGTATTAGAGACA GGAAGCATCAAGCCTGGGTAAATTGGAGGATCCAACCAAAGGTGCCACACCCAAAGTG GTGGAAAAATCGCTGAATATAACGCCAAATCCCACCATGTTGCCTGGGAGATCAGG GACCGGCTGCTGGCAGAGCGGGGTGTGACAATGACACCGTGCCTAGCGTCAGTCCATC AACAGGATCATCCGGACAAAAGTACAGCAGCCACCAACCAACAGTCCCAGCTCCAGT CACAGCATAGTGTCCACTGGCTCCGTGACGCAGGTGTCTCGGTGAGCACGGATTGGCC GGCTCGTCGTACTCCATCAGCGCATCCTGGCATCACGTCCCCAGCGCCACACCAAC AAGCGCAAGAGAGACGAAGGTATTCAAGGAGTCTCCGGTGCCAACGGCCACTCGCTTCCG GGCAGAGACTTCCCTCGGAAGCAGATGCGGGAGACTTGTTCACACAGCAGCAGCTGGAG GTGCTGGACCGCGTGTGAGAGGGCAGCACTACTCAGACATCTCACCACCAAGAGGCC ATCAAGCCCAGCACACAGAGTATTAGCCATGGCTCGCTGGCTGGTGGCTGGAC GACATGAAGGCAATCTGCCAGCCCCACCCCTGCTGACATGGGAGCAGTGTGCCAGGC CCGAGTCCTACCCATTGTGACAGGCCGTGACTTGGCAGCAGACCCCTCCCCGGGTAC CCTCCACACGTCCCCCGCTGGACAGGGCAGCTACTCAGCACCGACGCTGACAGGGATG GTGCTGGAGTGTGAGTTCCGGAGTCCCTACAGCCACCCCTCAGTATTCTCGTACAAC GAECTCTGGAGGTTCCCAACCCGGGCTGCTTGGCTCCCCCTACTATTAGCGCTGCC GCCGAGGAGCCGCCACCTGACGCCACTGCCTATGACCGTCACTGACCCCTGGAG CCAGGGGGCACAAACACTGATGGCACCTATTGAGGGTGAAGGCCACCCAGCCCTCCTG AAAGATGCCAGAGAGCCCAGTGGACCGTCCCCCAGCATCCCCACTGCTGAAGCTCCC CTCTCCTCTCTCCAGGGACTCTGGGCCCTTGGTGGGGCCGGTGGACTCTGGA TGCTTGCTATTCTAAAGCAATCTATGAGCTTCTCCGATGCCACTGGTCTCTGC AAACCAATAGACTGTCTGCAAATAACCGCAGCCCCAGCCAGCCTGCTGCCAGCTCCAGC TGTCTGACTATCCATCCATCATAACCAACCCCCAGCCTGGGAAGGGAGAGCTGCTTTGTTG CTTCAGCAGCACCCATGTAATACCTTCTGCTTTCTGTTGGCAGGTGCCTGAGAGCTGCT GAAGACTGCTCCACCCATGATGCATCTCGCACTCTGGTGCATACCGGACATCTAGAC CTATGGCAGAGCATCCTCTGCCCTGGGTACCCCTGGCAGGTGCCTGAGAGCTGCT CAAGATGGAGGATGCTGCCCTGGGCCCCAGCCTCTGCTCATCCCTCTTCTTGTAT CTTACGAGGAGTCTCACTGGCTGGTGTGCTGCAGGCTCCCCCTGAGGGCCCTCTCCA AGAGGAGCACACTTGGGAGATGCTGGTCTGCCCTGCCATTCTGGGACCGATG CAGTATCAGCAGCTTTCCAGATCAAAGAACTCAAAGAAAAGTGTCTGGAGATTCC CAGCTACTTTCCGAAGCAGAATGTCATCCGAGGTATTGATTACATTGTGGACTTGAAT GTGAGGGCTGGATGGGACGCGAGGAGTATCTGATCCAGCCAAGGAGGGGCTGAGGCT CTCCCTACTCCCTCAGCCCCCTGGAACGGTGTGTTCTGAGGCATGCCAGGTTCAAGTCAC

10

			TTGGACACCTGCCATGGACACTTCACCCACCCCTCAGGACCCCAGCAAGTGGATTCTGG GCAAGCTGTCGCGGTGATGTAGACAATAATTAAACACAGAGGACTTCCCCCACACCCAG ATCACAAACAGCCTACAGCCAGAACCTCTGAGCATCCTCTCGGGGCAGACCCCTCCCCGTC CTCGTGGAGCTTAGCAGGCAGCTGGGCATGGAGGTGCTGGGCAGATGCCTAAT TTCGCACAATGCATGCCACCTGTTGATCTAAGGGGCCGATGGTCAGGGCACGGCCA AGGGCCACGGAACTTGGAGAGGGAGCTTGGAGAACTCACTGTGGCTAGGGTGGTCAGA GGAAGCCAGCAGGGAAAGATCTGGGGACAGAGGAAGGCCCTCTGAGGGAGGGCAGGAGA GCAGTGAGGAGCTGTGACCTGGAGTGAATTGACATGGGGTGCAGGTGCCAT CATCTCTTACCTGGGCCTTAATTCTTGATAGTCTCTTGTCAAGTCAGAACAGCC AGGTAGAGCCCTGTCCAAACCTGGCTGAATGACAGTGTAGAGAGGGGCTTGGCCTTC TTAGGTGACAATGTCCCCATATCTGTATGTCAACCAGGATGGCAGAGAGGCCAGGGCAGA AGAGACTGGACTTGGATCAGCAGGCCAGCAGGTCTGTCTGGCCATGTC TTGCTGTGGACCTCAGACAAAACCTGCACCTCTTGAGCCTGGCTGCCTGGTGCA GCAGGGCATCTGTAGGCCACCCACAGCTTCTCCCTCTCTCCAGGGCCTTCAGTGGTGCCAGA CCAGGCCTGGGAGGATAACAGCCCTCCACCAGAAGAAGCTATGCCCTTGAGTTGACCA CCTGCCAGAGAGGCTGCAGTCACCTCTTACAGTCCAGACCTTGCTCCAGGAGGC CCAGGTGCTGCAGGGGCTCCCGAGCTCTCCCGAGGAGGCCAACGCCCTAGGCCCTGCA AAGGCTGGCTCCAGAGGAGGCTACAGCCCTCCCCCTGAGGAGACTATGCCATTGAGCTT GATGGAGAAGGATTGGGACGACAGCCCACCCCGGGCTTCCCGAGTTATCGCACAA GTCGACGGCAGCAGCCAGTCGCGCAGTCGCGCCTCGAGTGCAGTCCGCCTCACTCCC GCCCGAACGCGCCTCCCTCTGGTCCAGCGCCATGGCAGCCATCCAAAGAGGCT GTCAAGACCTCTTAACCTCACGGCAGCAGCCCTGGATGGAGATCTCCGGACCCCC TTCGAGATTGGCAGCGCCCCGCTGGGTGACGACACTCCGTCAACATGGACAGCCC CCAATCGCCTGACGGCCGCCATCAAGGCTCCGGAGGCCAGATAAGAGAGAGCGA GCAGAGAGACCCCAGTTGAGGAGGAAGCAGCAGAGATGGAAGGAGCGCTGATGCC GAGGGAGGAAAGTACCCCTCCGGGGTACGGATCCCTGCCGCCGGGCAGCCCTAGCG GATACCGCTGCCAGGGCAGCCCTGCAGCCCAGCCGATCCTGACTCCGGGCAACCC GAAGATCCCAGACTCCGGACAGCACCGCCGATCCTGACTCCGGGCACTCGCAGCCGAT CCGACTCCGGGGCAGCCCTGCCGCCAGCCGATCCGACTCCGGGCGGCCCTGAC GCCCGAGCCGATCCGACTCCGGGGCGCCCTGACGCCAGCCGATCCAGATGCC GCCGCCCTGAGGCTCCGCCGCCCTGCGCTGCTGAGACCCGGCAGCCCATGTCGCC CCAGCTGCCAGACGCAGGGCTCCACTGCCAGCCGCTCTGCCACCCGGCAGCC CAAGTCCGGGGCGGCCCTGCAGCCCTGCCCTCCGGGCCAGACGCAAGATCCATCTC AGACCCCCCAGCCCCGAGATCCAGGCTGCCGATCCGCTACTCCGGGCCACTCGC TCTGCCCTGGGGGCAAGTCCGAGAGCAGCCGGCCCGCGTACTACCATGAAGGG GTGCCAGCAGCAGATGACTCCAGCGAGACGAGTCCGACGATGGACCTCCGGATGC CTCCGCTGGTTTCAGCATGGCAAATGCCGCCGCCAAAGCCCCAGCGCAACTTACTC CGCAACTTCTCGTCAAGCCTCGGGGCTGCTCGTCAAGTCAAGGTAACCGCT AAAGCCTCGCCTCTCAAGGTCAAGAAGGTACCCCTGGCGAGAAGCGCAGACAGATG CGCAAAGAAGCCCTGGAGAAGCGGGCCAGAAGCGCAGAGAAGAAACGCAAGTC
	S000056	F37	164

			ATCGACAAACAACCTCCAGGACGAAAAGATGGGCTACATGTGTACGCACCGCCTGCTGCTT CTAG
S000058	F38	165	CTGCAGCTCTAGGACCCGGTTCTTTACTGATTAAAAACAAAACAAAAAAATAAA AAAGTTGTGCCTGAAATGAATCTGTTTTTTTATAAAGTAGCCGCCTGGTACTGTGT CCTGTAAAATACAGACATTGACCCCTGGTGTAGCTCTGTCACACTTATACACGGGAA TGGATGGGTCTGATTCTGGCCCTCTTCTGAATTGCCATATACAGGGCCCTGGCCA GTGGACTGAAGGCTTGCTAAGATGACAAGGGTCAGCTCAGGGATGTGGGGAGGGCG GTTTATCTCCCCCTTGTCTGTTGAGGTTTGTACTCTGGTAAGAGGGCGTTATCT TTGTAACACGAAACATTTTGTCTCCAGTTCTGTTAATGGCAGAACAGATGAAAG CGAATAAAAGTTTACTGATTGAGACACTAGCACCTAGCGCTTCAATTGAAACGT CCCGTGTGGAGGGCGGGCTGGGTGCGGCTGCCGATGACTCGTGGTCGGAGGCCA CGTGGCCGGGCGGGACTCAGGCCCTGGCAGCCGACTGATTACGTAGCGGGCGGGCC GGAAGTGCCTGCCTGGGTGTTATGGCGTCCGGGTCTCAACATTTTC CCGGTCTGTGGCTCAAATCTGTCCAAGCAGAGGCAGTGGAGCTGAGGTTCTGCTGG TGTGAAATGACTGAGTACAAACTGGTGGTGGAGCAGGTGGTGTGGAAAAGCGCA CTGACAATCCAGCTAATCCAGAACCACTTGTAGATGAATATGATCCCACCATAGAGGAT TCTTACAGAAAACAAGTGGTTAGATGGTAAACCTGTTGGACATACTGGATACA GCTGGACAAGAAGAGTACAGTCCATGAGAGACCAATACATGAGGACAGGCGAAGGCTTC CTCTGTGTATTCGCATCAATAAGCAAGTCATTGCGGATATTACCTCTACAGGGAG CAGATTAAGCGAGTAAAGACTCGGATGATGTACCTATGGTGTAGTGGAAAACAAGTGT GATTGCCAACAAGGACAGTTGATACAAACAAGCCCACGAACTGGCCAAGAGTACGGG ATTCCATTCTTGAACCTCAGCCAAGACCAGACAGGGTGTGAAGATGTTTACACA CTGGTAAGAGAAATACGCCAGTACCGAATGAAAAAAACTCAACAGCAGTGTGATGGACT CAGGGTTGTATGGATTGCCATGTGTGGTGTGAAAGATACTTTAAAGTTTGTCA GAAAAGAGCCACTTCAAGCTGCACTGACACCCCTGGCTGACTTCCCTGGAGGAGAAGTA TTCTGTTGCTGCTTCAGTCTCACAGAGAAGCTCTGCTACTTCCCTGGCTCTAGTAG TTTACATAATACTCTTGTGAGAAGTTCTCAGAATAACTACCTCCTACTGGCTGT CTGACCAGAGAATGCACCTCTGTTACTCCCTGTTATTTCTGCCCTGGTTCTCCAC AGCACAAACACACCTCAACACACCTGCCACCCAGGTTTCTGAAAGCAGTTC ATGCTGAAACAGAGAACCAACCGCAAACGTGAAATTCTATTGAAAACAGTGTCTTGAG CTCTAAAGTAGCAACTGCTGGTGTGTTTACTGTTGAACTTAGAACTAT GCCTAATTTGGAGAAATGTCATAAATTACTGTTGCCAGAAATAGTTATTGTC TGTTGGTTGTATGTTATGTTATCGGCTCTTCTCAAACCTGGCATCTGCTCTAGATT CATAAATACAAAAATGAATACTGAATTGAGTCTATCCTAGTCTCACAACCTTGACGT AATTAATCCTAGTGGAAATAGATGTCATAAAATCCTATGCAATGAAATGATGCTGA GATACGTCTGACTTATCACCATTGAAGGAAAGCTATATCTATTGAGAGCAGATGCC ATTTGTACATGTGAAATTGGTTCCAGAGGGCTGTTGGGGCTTCCAGGAGAA AGATGAAACAGAGAACATGAATAATTCACTTAAATTTACCTAATCTCCACTTT TTTCATAGGTTACTACCTATACAATGTGTAATTGTTCCCTAGCTACTGATAAAC CTAATATTCAATGAACCTCATTGTTCAAAATTGTCATACCAAGAACGCTACAT TTGCAGATGTCATAATATTGAAAACCTTGGTGCATTGTTATTAAATAGCTGTGATCAGT GATTTCAAACCTCAAATATAGTATATTAAACAAATT
S000072	F39	166	TTGGAGCTGCCGCCGGGACTCCGTCCAGCAGGACATGGATTGATTGACATACTT TGGAGGCAAGATATAGATCTGGAGTAAGTCAGGAGAAGTATTGACTTCAGTCAGCGACGG AAAGAGTATGAGCTGGAAAAACAGAAAAAAACTGAAAAGGAAAGACAAGAACAACTCCAA AAGGAGCAAGAGAAAGCCCTTCACTCAGTACAACTAGATGAAGAGACAGGTGAATT CTCCCAATTAGCCAGCCAGCACACCCAGTCAGAAACCAAGTGGATCTGCCAACTACTCC CAGGGTGCCTCACATTCCAAATCAGATGCTTGTACTTGTGACTGCATGCAGCTTGT

			GCGCAGACATCCCGTTGTAGATGACAATGAGGTTCTCGGCTACGTTCAGTCACTT GTTCTGATATTCCCGTCACATCGAGAGCCCAGTCCTCATGGCTACTAATCAGGCTAG TCACCTGAAACTCTGTTGCTCAGGTAGCCCCGTTGATTAGACGGTATGCAACAGGAC ATTGAGCAAGTTGGGAGGAGCTATTATCCATTCTGAGTTACAGTGTCTTAATATTGAA AATGACAAGCTGGTTGAGACTACCAGGTTCCAAGTCCAGAAGCCAAACTGACAGAAGTT GACAATTATCATTTTACTCATCTACCCCAATGGAAAAAGAAGTAGGTAACGTAGT CCACATTTCTTAATGCTTGAGGATTCTCAGCAGCATCCTCTCCACAGAACAGACCC AACCAAGTTGACAGTGAACCTCATAAATTAGATGCCACAGTCACACAGATTTGGTGA GAATTTATTCTGCTTCAGCTGAGCCCAGTACAGCAACAGCATGCCCTCACCTGCT ACTTAAAGCCATTCACTCTGAACCTCTAAATGGGCCATTGATGTTCTGATCTATCA CTTGCAAAGCTTCAACAAAACCACCCCTGAAAGCACAGCAGAATTCAATGATTCTGAC TCCGGCATTCACTAAACACAAGTCCCAGTGGCATCACCAGAACACTCAGTGGAACT TCCAGCTATGGAGACACACTACTTGGCCTCAGTGATTCTGAAGTGGAAAGAGCTAGATAGT GCCCTGGAAGTGTCAAACAGAAATGGCTCTAAACACCCAGTACATTCTCTGGGGATATG GTACAACCCCTGTCAACCCTCAGGGGCAGAGCACTCAGTGCATGATGCCAATGTGAG AACACACCAGAGAAAGAATTGCTGTAAGTCCTGGCATCGAAAACCCATTACAAAAA GACAACATTCAAGCCGTTGGAGGCTCATCTACAAGAGATGAACTTAGGGCAAAGCT CTCCATATCCCATTCCCTGAGAAAAAAATCATTAACCTCCCTGTTGACTTCAACGAA ATGATGTCAAAGAGCAGTTCAATGAAGCTCAACTTGCATTAATTGGGATATACGTAGG AGGGTAAGAATAAAAGTGGCTGCTCAGAATTGCAAGAAAAGAAAATGGAAAATATAGTA GAACAGAGCAAGATTAGATCATTGAAAGATGAAAAAGAAAATTGCTCAAAGAAAAA GGAGAAAATGACAAAGCCTTCACCTACTGAAAAACAAACTCAGCACCTTATCTGAA GTTTCAGCATGCTACGTGATGAAGATGGAAAACCTTATTCTCTTAGTGAATACTCCCTG CAGCAAACAAGAGATGGCAATGTTCTGTTCCAAAAGTAAGAACGCCAGATGTTAAG AAAAACTAGATTAGGAGGATTGACCTTTCTGAGCTAGTTTTTGACTATTAACT AAAAGCTCTACTGTGATGTGAAATGCTCATACTTATAAGTAATTCTATGCAAATCAT AGCCAAAACTAGTATAGAAAATAACGAAACTTAAAAGCATTGGAGTGTCACTGTT TGAATCACTGAGTTCACTTAACTGAAACATTCTAGGACACCATTGGCTAGTT CTGTGTAAGTGTAAATACTACAAAACCTTTACTGTTCTTATGTCATTGTTATAT TCATAGATTATATGATGATGACATCTGGCTAAAAGAAATTATTGCAAACAACTAACCA CGATGTACTTTTATAAAACTGTATGGACAAAAAATGGCATTTTATAATTAAATTG TTAGCTGGCAAAAAAAAAAATTGTTAAGAGCTGGTACTAATAAAGGATTATTATG ACTGTT
S000083	F40	167	GGGGGCAGAGGGAGCGAGCGGGCGGCCCTAGGGTGCAGAGGCCGGCGAGCAGAGTTG CGCTCGGGCGTCTGGAAAGGGAGTCCGGAGCCAACAGGGGGCTCGCCTCTGGCCCA GCCCTCCGGAGCCAACAGGGACTTCGCCTCTGGCCCAGCCCTCCCGCTGATCCCCCAG TCAGCGTCCGCAAGCCTTGCCGCATCCACGAAACTTGGCCACTTGCCACTGCGGGCGTACACT TTGCACTTGAACCTACAACACCCGAGCAAGGACCGACTCTCCGACGCGGGAGACTAT TCTGCCATTGGGACACTTCCCCCGCTGCCAGGACCCGGTTCTCTGGAAGGCTGTC CTTGAAGCTCTTAGACGCTGGAGTTTTGGAAAGTGGAAAGCAGCCTCCCGCAG ATGCCCTCAACGTTAGCTTACCAACAGGAACATGACCTCGACTACGACTCGTGCAG CCGATTCTACTGCGACGAGGAGGAGAACTTCTACCGAGCAGCAGCAGAGCGAGCTG CAGCCCCCGCGCCAGCGAGGATATCTGGAAGAAATTGAGCTGCTGCCCACCCCGCC CTGCCCCCTAGCCGCCGCTCCGGGCTCTGCTGCCCTCTACGTTGGCTCACACCCCTC TCCCTCGGGGAGACAACGACGGGGTGGCGGGAGCTCTCCACGCCGACCGAGCTGGAG ATGGTGACCGAGCTGCTGGGAGGAGACATGGTGAACCAGAGTTCTGCGACCCGGAC GACGAGACCTTCAAAACATCATCATCCAGGACTGTATGTTGAGCGGCTCTCGGCC GCCGCCAAGCTCGTCTCAGAGAAGCTGGCTCTACCAGGCTGCGCGCAAAGACAGCGGC AGCCCGAACCCCGCCGCCAGCGTCTGCCACCTCCAGCTTACCTGCAG

			CTGAGCGCCGCCGCCTCAGAGTCATCGACCCCTCGGTGGTCTTCCCTACCCCTCAAC GACAGCAGCTGCCAAGTCCTGCGCCTCGAAGACTCCAGCGCCTCTCCGCTCG GATTCTCTGCTCTCCCGACGGAGTCTCCCCGCAGGGCAGCCCCGAGCCCTGGTGCTC CATGAGGAGACACCGCCCACCACCGAGCAGCAGCAGTCTGAGGAGGAACAAGAAGATGAGGAA GAAATCGATTTGTTCTGTGGAAAAGAGGGCAGGCTCTGGCAAAGGTAGAGTCTGG TCACCTCTGCTGGAGGCCACAGCAAACCTCCTCACAGCCACTGGTCTCAAGAGGTGC CACGTCTCCACACATCAGCACAACACTACGCAGGCCCTCCACTCGGAAGGACTATCCT GCTGCCAAGAGGGTCAAGTGGACAGTGTAGACAGTCTGAGACAGATCAGCAACAACCGA AAATGCACCAGCCCCAGGTCTCGGACACCGAGGAGAATGTCAAGAGGCGAACACACAAC GTCTGGAGGCCAGAGGAGGAACGGAGCTAAAACGGAGCTTTTGCCTCGTGACAG ATCCCGGAGTTGAAAACATGAAAAGGCCCAAGGTAGTTATCCTAAAAAGCCACA GCATACATCCTGTCCGTCCAAGCAGAGGAGCAAAGCTCATTCTGAAGAGGACTTGTG CGGAAACGACGAGAACAGTGAACACAAACTGAACAGCTACGGAACTCTGTGCGTAA
S000087	F41	168	GGGGCGAGAGGGAGCGAGCGGGCGGCCCTAGGGTGAAGAGCCGGCGAGCAGAGTTG CGCTGGGGCGTCTGGAAAGGGAGTCTGGAGCCAACAGGGGCTTCGCCCTGGCCCA GCCCTTCCGGAGCCAACAGGGACTTCGCCTCTGGCCAGCCCTCCGCTGATCCCCAG TCAGCGGTCCGCAAGCCTGCCGCATCCACGAAACTTGGCCACTGCGGGCTACACT TTGCACTGAACTACAACACCCGAGCAAGGACCGACTCTCCGACGCCGGAGACTAT TCTGCCCATTGGGACACTTCCCGCCGCTGCCAGGACCCGGTTCTGGAAAGGCTGTC CTTGAAGCTCCTTAGACGCTGGAGTTTCTGGGAAGTGGAAAGCAGCCTCCGCGACG ATGCCCTCAAGTTAGCTTACCAACAGGAACATGACCTCGACTACGACTCGTGAG CCGTATTTCTACTGCGACGAGGAGGAGAACCTTACCAAGCAGCAGCAGAGCGAGCTG CAGCCCCCGGCCAGCGAGGATATCTGGAAAGAAATTGAGCTGCTGCCCACCCGCC CTGCCCCCTAGCCGCCCTCCGGCTGCTGCCCTCTACGTTGGTGTACACCCCTC TCCCTCGGGAGACAACGACGGGGTGGCGGGAGCTTCCACGCCGACAGCTGGAG ATGGTACCGAGCTGCTGGAGGAGACATGGTGAACCAGAGTTCATCTGCGACCCGGAC GACGAGACCTCATAAAAACATCATCCAGGACTGTATGTGGAGGGCTTCTGGCC GCCGCAAGCTCGTCTAGAGAAAGCTGGCTTACCAAGGCTGCGCGCAAAGACAGCGGC AGCCCGAACCCGCCGCCAGCGTCTGCTCCACCTCCAGCTTGTACCTGAGGAT CTGAGCGCCGCCGCCTCAGAGTCATCGACCCCTGGTGTCTCCCTACCCCTCAAC GACAGCAGCTGCCAAGTCCTGCGCCTCGAAGACTCCAGCGCCTCTCCGCTCG GATTCTCTGCTCTCCCGACGGAGTCTCCCCGCAGGGCAGCCCCGAGCCCTGGTGCTC CATGAGGAGACACCGCCCACCACCGAGCAGCAGTCTGAGGAGGAACAAGAAGATGAGGAA GAAATCGATTTGTTCTGTGGAAAAGAGGGCAGGCTCTGGCAAAGGTAGAGTCTGG TCACCTCTGCTGGAGGCCACAGCAAACCTCCTCACAGCCACTGGTCTCAAGAGGTGC CACGTCTCCACACATCAGCACAACACTACGCAGGCCCTCCACTCGGAAGGACTATCCT GCTGCCAAGAGGGTCAAGTGGACAGTGTAGACAGTCTGAGACAGATCAGCAACAACCGA AAATGCACCAGCCCCAGGTCTCGGACACCGAGGAGAATGTCAAGAGGCGAACACACAAC GTCTGGAGGCCAGAGGAGGAACGGAGCTAAAACGGAGCTTTTGCCTCGTGACAG ATCCCGGAGTTGAAAACATGAAAAGGCCCAAGGTAGTTATCCTAAAAAGCCACA GCATACATCCTGTCCGTCCAAGCAGAGGAGCAAAGCTCATTCTGAAGAGGACTTGTG CGGAAACGACGAGAACAGTGAACACAAACTGAACAGCTACGGAACTCTGTGCGTAA

15

			GGAAAAGTAAGGAAAACGATTCCCTAACAGAAATGTCCCTGAGCAATCACCTATGAAC TGTTTCAAATGCATGATCAAATGCAACCTACAACCTGGCTGAGTCTTGAGACTGAAAG ATTTAGCCATAATGTAACACTGCCCAAATTGGACTTGGGCATAAAAGAACTTTTATGC TTACCATCTTTTTCTTTAACAGATTGTATTAAGAATTGTTTAAAAAATTT AAGATTACACAATGTTCTGTAAATATTGCCATTAAATGTAATAACTTTAATAAAA ACGTTTATAGCAGTTACACAGAATTCAATCCTAGTATATAGTACCTAGTATTATAGGTA CTATAAACCTAATTTTTATTAAGTACATTGCTTTAAAGTTGATT
S000090	F42	169	GGGGGCAGAGGGAGCGAGCGGGGGCGCTAGGGTGCAGAGGCGGGCGAGCAGAGTTG CGCTGCGGGCGTCCCTGGGAAGGGAGTCCGGAGCCAACAGGGGGCTCGCCTCTGGCCCA GCCCTTCCGGAGCCAACAGGGACTTCGCCTCTGGCCCAGCCCTCCCGCTGATCCCCCAG TCAGCGGTCCGCAAGCCTTGCCGCATCCACGAAACTTGGCCATACTGCGGGCGTACACT TTGCACTGAACTACAACACCCGAGCAAGGACCGACTCTCCGACGCGGGGAGACTAT TCTGCCCATTTGGGGACACTTCCCCGCCGCTGCCAGGACCCGGTTCTCTGGAAAGGCTGTC CTTGAAGCTCCTAGACGCTGGAGTTTCTGGGAAGTGGAAAGCAGCCTCCCGCAGC ATGCCCTCAACGTTAGCTTACCAACAGGAACATGACCTCGACTACGACTCGTGCAG CCGTATTTCTACTGCGACGGAGGAGAACTTCTACCAGCAGCAGCAGCAGAGCGAGCTG CAGCCCCCGGCCAGCGAGGATATCTGGAAAGAAATTGAGCTGCTGCCCACCCCGCCC CTGCCCCTAGCCGCCGCTCCGGCTCTGCTGCCCTCACGTTGGGTACACCCCTC TCCCTCGGGGAGACAACGACGGGGTGGCAGGAGCTTCTCCACGCCGACCGAGCTGGAG ATGGTACCGAGCTGGAGGAGACATGGTAACCAGAGTTCATCTGCGACCCGGAC GACGAGACCTCATAAAAACATCATCCAGGACTGTATGTGGAGCGGCTCTCGGCC GCCGCAAGCTCGTCTAGAGAAGCTGGCTCTACCAGGCTGCGCAGAACAGCGGC AGCCCGAACCCGCCGCGGCCAGCGTCTGCTCCACCTCCAGCTGTACCTGCAGGAT CTGAGCGCCGCCGCTCAGAGTGCATGACCCCTCGTGGCTTCCCTACCCCTCAAC GACAGCAGCTGCCAAGTCCGCTCGCAAGACTCCAGCGCCTCTCCGCTCTCG GATTCTCTGCTCTCCCGACGGAGTCTCCCGCAGGGCAGCCCCGAGCCCTGGTCTC CATGAGGAGACACGCCACCACGAGCAGCAGCAGCTGAGGAGGAACAAGAAGATGAGGAA GAAATCGATTTGTTCTGTGGAAAGAGGCAGGCTCTGGCAAAGGTCAAGAGTCTGG TCACCTTCTGCTGGAGGCCACAGCAAACCTCTCACAGCCCAGTGGCTCTAAGAGGTGC CACGTCTCCACACATCAGCACAACTACGCAAGCGCTCCCTCCACTCGGAAGGACTATCT GCTGCCAAGAGGGTCAAGTGGACAGTGTAGCTGAGACAGATCAGCAACACCGA AAATGCACCAGCCCCAGGTCTCGGACACCGAGGAGAATGTCAGAGGCGAACACACAA GTCTGGAGCGCCAGAGGAGGAACGAGCTAAACAGGAGCTTTTGCCTCGTGACCA ATCCCGAGTTGAAAACAATGAAAAGCCCCAAGGTAGTTATCCTAAAAAGGCCACA GCATACATCCTGCGTCCAAGCAGAGGAGCAAAGCTCATTCTGAAGAGGACTGTTG CGGAAACGACGAGAACAGTTGAAACACAAACTGAAACAGCTACGGAACACTTGTGCGTAA GGAAAAGTAAGGAAAACGATTCTTAACAGAAATGTCCCTGAGCAATCACCTATGAAC TGTTCAAATGCATGATCAAATGCAACCTACAACCTGGCTGAGTCTTGAGACTGAAAG ATTTAGCCATAATGTAACACTGCCCAAATTGGACTTGGGCATAAAAGAACTTTTATGC TTACCATCTTTTTCTTTAACAGATTGTATTAAGAATTGTTTAAAAAATTT AAGATTACACAATGTTCTGTAAATATTGCCATTAAATGTAATAACTTTAATAAAA ACGTTTATAGCAGTTACACAGAATTCAATCCTAGTATATAGTACCTAGTATTATAGGTA CTATAAACCTAATTTTTATTAAGTACATTGCTTTAAAGTTGATT
S000098	F43	170	TCGGAGACCACATTGCCCTGTCACACTATCCTACCAAGAAGAAATCTATTGTTT AGCCTGAGACACTCTTGAGGTAAAAAATTAGAATGAAAGAACCTTGGATGGTGAATGT GGCAAAGCAGTGGTACCAACAGCAGGAGCTTGGACAAAATTAAAGAAGAACAGACAAT GCTCAAGAGTATGGATGTGTCACAGCCAAAACCTCAAGAAAGTAAATTGAAATTGGT GGTGTGTTCTAGTTAATGAGAGACCTATTGCCAGCAGTTGAACCCAGGCTTCAGCTT TCTTTGCATCATCGGCCAAGTGTGTTGCTTCCCTCAGTCCAGCTGTTGCTATTAAG

			GTTTTTGTCTGGTTGAAAAAAATGCTTATAAGGCCAAACTGCATATCATAAGACA GGATCTACTCAGCTCTGCTCCACACGATGCATCACAGACATTCTCACCTGCC CTGCCACCTCCCTCCAAGAAAACCTGCACAAACTGCTCGAAAGACATTAAATCCTAAG GATGTGATCACAACCTCGCTTGAGAATTCTATCCTAGCAAAGATTCTGCAGCCAATCA TGCTGTATCTTATGAGCTAAAGAAAAACCTGTTACCATATACCAAAAGCATT TCAACTAAGTGCAGTATGTCAGAAGAATGCTGATACCGATTGAAGTAAATATCAA AATGTGGTACATGGTCTTGTAGTGCTGTTTCAAATTCACTCTACAAACAAAC CTCACCATGAACGTGGTGAGAACTGTGGAGCTATTGCTATAGTAGCTCTGGTCTTGC CAATCCCAGAAGGTTTGTCAACAGTGCACGGCATAACAAGCAGAATTCTGCC ATTCCCTCCATATGCCCTGGGAAGTCATTGAGGCCCTCAGCTGAAATGATTGAGACTACA AATGATTAGGAAAAACAGAGCTTCTGCTCTATTAAATTGCTTATCTGCTTACAGAGTT AAGACTGTTACTCTTCAGGTGTCAGGTTATGTCATAGTTGAACCTCAGCAATC CCTCAGTATCACCTAGCCATGTCAAATGGAACATATACAGCTCTGCAGCTCCAGTTGT GTGGTTGCTTCCAGAATGTTAGCAAGCAGGAAACAAACTCTCGGCGGTGCC CTGTCAGGGCCAAGTGGTTGAAGCCCCGCCCTCCTCCAGGTAGCAGGTCAATAGGA GGAGGTAACACCTCTGCCGTTCCCCAGCTCCATCCGTGGCTCTGCTGCAGCCAGCCTC CAACCTCTGGTGAACAATCCCAGCAAGTTGCTTAACCCATACAGTTAAACTCAAG TGTCACTGTAACCATCTATTGCCACAAAACCAGAACATTCTTTTACAAGGGTAA ATGTTCTGTTTGTGGCAAGAATTGCTCTGATGAATACAAGAAGAAAAAAATGTTGT GCAATGTGTGACTACTGTAACCTGCAGAAAATTATAAAGGAGACTGTGCGATTCTCAGGG GTTGATAAGCATTCTGTAGTGAAAGTTGCAAATTCTCTGCCGTGACTTGGAGAA CGATGGGAAACTACTGTAAGATGTGAGCTACTGTTCACAGACATCCCCAAATTGGTA AAAAATGATTGGAGGGCAAGTTAGAAGAGTTGTTGAAGATTGATGTCCAAATT ACAGTTCTGTTTATCAGATGGCAAGTGTGATGGTTGAAACGACAGGGTAAACTAAGC GAGTCCATAAAGTGGCGAGGCAACATTAACATTCTGTAACCTATTGTCAGG TTTGTCATCAGCAAATTATGAATGACTGTCTCCACAAAATAAGTAAATATTCTAA GCAAAACTGCTGTGACGGAGCTCCCTCTGCAAGGACAGATAACACACCAGTTAA AGTGTGATGTCATTGGCAAAATACCTGCTACCTTATCTACAGGGAACACTAACAGTGT TTAAAGGTGAGTTACTAAAGAGGCAGCAAAGATCATTCAAGATGAAAGTACACAGGAA GATGCTATGAAATTCCATCTTCCAACTTCCCAGCCTCCAGGCTTTAAAGAACAAA GGCATATCATGCAAACCGTCACACAGACCAAGGCCACTTCTGCAAACACATACACAG CACAAAGAATGTCAGACAGAAATGCCCTGTCAGTTGCTGAGGTGTTCCCGCTGAA GTATTGGTACCCAGCAACTCCCTGAACCAATAGCTGTGGCTTCTGGAACGT CTGGCTGGTTGCTGGTACCGGCCACCGATGCTGCAAGAGATCAGTAAC
S000104	F44	171	GGGGGCAGAGGGAGCGAGCGGGCGCCTAGGGTGCAGAGGCCGGCGAGCAGAGTTG CGCTGCGGGCGTCTGGAGGGAGTCCGGAGCCAACAGGGGGCTCGCCTCTGGCCA GCCCTCCGGAGCCAACAGGGACTTCGCTCTGGCCAGCCCTCCCGCTGATCCCCAG TCAGCGGTCCGCAAGCCTTGCAGCATAACAGGAAACTTGCCTACTGCGGTACACT TTGCACTGAACTTACAACACCCGAGCAAGGACGCGACTCTCCGACGCGGGGAGACTAT TCTGCCATTGGGACACTTCCCCGCCGCTGCCAGGACCCGGTTCTGGAAGGCTGTC CTTGAAGCTCTAGACGCTGGAGTTTGGAGAAGTGGAAAGCAGCCTCCCGCAG ATGCCCTCAACGTTAGCTTACCAACAGGAACATGACCTCGACTACGACTCGGTGAG CCGTATTCTACTGCGACGAGGAGGAGAACCTTCTACCGCAGCAGCAGCAGAGCTG CAGCCCCCGGCCAGCGAGGATATCTGGAAGAAATTGAGCTGCTGCCACCCCGCCC CTGCCCCCTAGCCGCCGCTCCGGCTGCTGCCCTCTACGTTGCGGTACACCCCTC TCCCTCGGGAGACAACGACGGCGGTGGAGCTTCTCACGCCGACCGAGCTGGAG ATGGTACCGAGCTGCTGGAGGAGACATGGTAACCAAGAGTTCATCTGCGACCCGGAC GACGAGACCTTACCAAAACATCATCCAGGACTGTATGTTGGAGCGGTCTCGGCC GCCGCCAAGCTCGTCTAGAGAAGCTGGCCTCTACCGGCTGCGCGCAAAGACAGCGGC

			AGCCCGAACCCGCCGGCACAGCGTCTGCTCACCTCCAGCTGTACCTGCAGGAT CTGAGCGCCGCCGCTCAGAGTCATCGACCCCTCGTGGCTTCCCCTACCCCTCAAC GACAGCAGCTGCCAAGTCCTCGCCCTCGCAAGACTCCAGCGCCTCTCTCCGCTCG GATTCTCTGCTCTCTCGACGGAGTCCTCCCCGAGGGCAGCCCCGAGCCCCCTGGTGCTC CATGAGGAGACACCGCCACCACCAAGCAGCGACTCTGAGGAGGAACAAGAAGATGAGGAA GAAATCGATTTGTTCTGTGGAAAAGAGGCAGGCTCCTGGCAAAAGGTCAAGAGTC TCACCTTCTGCTGGAGGCCACAGCAAACCTCCTCACAGCCCACGGCTCTCAAGAGGTGC CACGTCTCCACACATCAGCACAACACTACGCAGCGCCTCCACTCGGAAGGACTATCCT GCTGCCAAGAGGGTCAAGTGGACAGTGTCAAGTCAGACAGATCAGCAACAACCGA AAATGCACCAGCCCCAGGTCTCGACACCGAGGAGAATGTCAAGAGGCAGACACACAAC GTCTGGAGCGCCAGAGGGAGAACGAGCTAAACGGAGCTTTTGCCCTGCGTACCGAG ATCCCGGAGTTGAAAACAATGAAAAGCCCCAAGGTAGTTATCCTAAAAAGCCACA GCATACATCCTGTCCGTCCAAGCAGAGGAGCAAAGCTCATTCTGAAGAGGACTTGTG
S000106	F45	172	GGGGGCAGAGGGAGCGAGCGGGGGCCGCTAGGGTGCAGAGGCCGGCGAGCAGAGTTG CGCTCGGGCGTCCTGGAAAGGGAGTTCCGGAGCCAACAGGGGGCTCGCCTCTGGCCCA GCCCTCCGGAGCCAACAGGGGACTTCGCTCTGGCCAGGCCCTCCCGCTGATCCCCAG TCAGCGGTCCGCAAGCCTTGCCGCATCCACGAAACTTGCCTACTGCAGGGCTACACT TTGCACTGAACCTACAACACCCGAGCAAGGACCGACTCTCCGACGCCGGAGACTAT TCTGCCCTTTGGGACACTTCCCGCCGCTGCCAGGACCCGGTTCTCTGGAAAGGCTGTC CTTGAAGCTCCTAGACCGTGGAGTTTCTGGAAAGTGGAAAGCAGCCTCCCGACG ATGCCCTCAACGTTAGCTTCACCAACAGGAACATGACCTCGACTACGACTCGTGCAG CCGTTACTGCAGGAGGAGGAACCTTCTACCAGCAGCAGCAGCAGAGCGAGCTG CAGCCCCCGGCCAGCGAGGATATCTGGAAAGAAATTGAGCTGCTGCCCACCCGCC CTGCCCCTAGCCGCCGCTCGGGCTCTGCTGCCCTCACGTTGGTACACCCCTTC TCCCTCGGGAGACAACGACGGGGTGGCGGGAGCTTCTCACGCCGACCGAGCTGGAG ATGGTGACCGAGCTGGAGGAGACATGGTGAACCAGAGTTTATCTGCGACCCGGAC GACGAGACCTTCATAAAAACATCATCATCCAGGACTGTATGAGGGCTCTCGGCC GCCGCAAGCTCGTCTCAGAGAAAGCTGGCTCTACCAGGCTGCGCAGACAGCGGC AGCCCGAACCCGCCGGCCACAGCGTCTGCCACCTCCAGCTGTACCTGCAGGAT CTGAGCGCCGCCGCTCAGAGTCATCGACCCCTCGTGGCTTCCCCTACCCCTCAAC GACACCGACTCGCCCAAGTCCTGCCTCGCAAGACTCCAGCGCCTCTCTCCGCTCG GATTCTCTGCTCTCTCGACGGAGTCCTCCCCGAGGGCAGCCCCGAGCCCCCTGGTGCTC CATGAGGAGACACCGCCACCACCAAGCAGCGACTCTGAGGAGGAACAAGAAGATGAGGAA GAAATCGATTTGTTCTGTGGAAAAGAGGCAGGCTCCTGGCAAAAGGTCAAGAGTC TCACCTCTGCTGGAGGCCACAGCAAACCTCCTCACAGCCCACGGCTCTCAAGAGGTGC CACGTCTCCACACATCAGCACAACACTACGCAGCGCCTCCACTCGGAAGGACTATCCT GCTGCCAAGAGGGTCAAGTGGACAGTGTCAAGTCAGACAGATCAGCAACAACCGA AAATGCACCAGCCCCAGGTCTCGACACCGAGGAGAATGTCAAGAGGCAGACACACAAC GTCTGGAGCGCCAGAGGGAGAACGAGCTAAACGGAGCTTTTGCCCTGCGTACCGAG ATCCCGGAGTTGAAAACAATGAAAAGCCCCAAGGTAGTTATCCTAAAAAGCCACA GCATACATCCTGTCCGTCCAAGCAGAGGAGCAAAGCTCATTCTGAAGAGGACTTGTG

			CGGAAACGACGAGAACAGTTGAAACACAAACTGAACAGCTACGGAACCTTGCGTAA GGAAAAGTAAGGAAAACGATTCCCTCAACAGAAATGCTTGAGCAATCACCTATGAAC TGTTCAAATGCATGATCAAATGCAACCTCACAACCTGGCTGAGTCTTGAGACTGAAAG ATTTAGCCATAATGTAAACTGCCTCAAATTGGACTTGGCATAAAAGAACCTTTATGC TTACCATTTTTTTCTTAACAGATTGTATTAAAGAATTGTTTTAAAAAATT AAGATTACACAATGTTCTGTAAATATTGCCATTAATGTAATAACCTTAATAAAA ACGTTATAGCAGTTACACAGAATTCAATCCTAGTATATAGTACCTAGTATTATAGGTA CTATAAACCTAATTTTTATTAAAGTACATTTGCTTTAAAGTTGATT	
S000107	F46	173	GGGGCGAGAGGGAGCGAGCGGGCCGCCTAGGGTGCAGAGGCCGGCGAGCAGAGTTG CGCTCGGGCGTCCTGGAGGGAGTCCGGAGCCAACAGGGGCTTCGCCCTGGCCCA GCCCTCCGGAGCCAACAGGGGACTTCGCCTCTGGCCCAGCCCTCCGCTGATCCCCAG TCAGCGGTCCGCAAGCCTGCGCATCCACGAAACTTGCCTACTGCGGGCTACACT TTGCACTTGAACCTACAACACCCGAGCAAGGACCGACTCTCCGACGCCGGAGACTAT TCTGCCCATTTGGGACACTTCCCGCCGCTGCCAGGACCCGGTCTCTGGAAAGGCTGTC CTTGAAGCTCCTTAGACGCTGGAGTTTCGGGAAGTGGAAAGCAGCCTCCCGGACG ATGCCCTCAACGTTAGCTTCACCAACAGGAACATGACCTGACTACGACTCGGTGCA CCGTTCTACTGCGACGGAGGAGAACCTTACCAAGCAGCAGCAGCAGAGCGAGCTG CAGCCCCCGGCCAGCGAGGATATCTGGAAAGAAATTGAGCTGCTGCCACCCGCC CTGCCCCTAGCCGCCGCTCGGGCTCTGCTGCCCTCACGTTGGCTCACCCCT TCCCTCGGGGAGACAACGACGGCGTGGCGGGAGCTTCCACGGCCGACCAGCTGGAG ATGGTACCGAGCTGGAGGAGACATGGTAACCAGAGTTCATCTGCGACCCGGAC GACGAGACCTTCATAAAAACATCATCATCCAGGACTGTATGTTGGAGCGCTCTGGCC GCCGCCAAGCTCGTCTCAGAGAACGCTGGCTTACCAAGGCTGCGCAAAGACAGCGGC AGCCGAACCCGCCGCCAGCGGCCACAGCGTCTGCTCCACCTCCAGCTGTACCTGCAGGAT CTGAGCGCCGCCGCTCAGAGTCATCGACCCCTCGGTGGCTTCCCTACCCCTCAAC GACAGCAGCTGCCAAGTCTGCGCTCGCAAGACTCCAGCGCTTCTCCCGCTCG GATTCTGCTCTCTCGACGGAGTCTCCCGCAGGGCAGCCCCGAGCCCTGGTGTCT CATGAGGAGACCCGCCACCAAGCAGCGACTCTGAGGAGGAACAAGAAGATGAGGAA GAAATCGATGTTCTGTGGAAAGAGGCAGGCTCTGGCAAAGGTCAAGAGTCTGGA TCACCTCTGCTGGAGGCCACAGCAAACCTCCTCACAGCCCCTGGCTCAAGAGGTGC CACGCTCCACACATCAGCACAACACTCGCAGGCCCTCCACTCGGAAGGACTATCCT GCTGCCAAGAGGGTCAAGTGGACAGTGTAGCTGAGACAGATCAGCAACACC AAATGCACCAGCCCCAGGTCTCGGACACCGAGGAGAACATGTCAGAGGGCGAACAC GTCTGGAGGCCAGAGGAGGAACGGACTAAACGGAGCTTTTGCCTCGTGACCAG ATCCCGAGTGGAAAACAATGAAAAGGCCCAAGGTAGTTATCCTAAAAAGGCCACA GCATACATCCTGCGTCCAAGCAGAGGAGCAAAGCTCATTCTGAAGAGGACTGTT CGGAAACGACGAGAACAGTTGAAACACAAACTGAACAGCTACGGAACCTTGCGTAA GGAAAAGTAAGGAAAACGATTCCCTCAACAGAAATGCTTGAGCAATCACCTATGAAC TGTTCAAATGCATGATCAAATGCAACCTCACAACCTGGCTGAGTCTTGAGACTGAAAG ATTTAGCCATAATGTAAACTGCCTCAAATTGGACTTGGCATAAAAGAACCTTTATGC TTACCATTTTTTTCTTAACAGATTGTATTAAAGAATTGTTTTAAAAAATT AAGATTACACAATGTTCTGTAAATATTGCCATTAATGTAATAACCTTAATAAAA ACGTTATAGCAGTTACACAGAATTCAATCCTAGTATATAGTACCTAGTATTATAGGTA CTATAAACCTAATTTTTATTAAAGTACATTTGCTTTAAAGTTGATT	
20	S000114	F47	174	GCATCCGGCATCTGACGTGGTTATGCTGCCAGTTGGCCGCACTGTAGGAAAG TAACCTCAGCTGCAAGCCCCAAAGCGAGTGAGCCGAGCCGGAGCCATGGAGGGCCAGAGCG TGGAGGAGCTGCTCGCAAAGGCAGAGCAGGAGGAGGAGAGAACGTTGCAACGCCATCACGG TGCACAAGGAGCTGGAGCTGCAGTTGACCTGGCAACCTGCTGGCGTCGGACCGAAC CCCCGACCGGGCTGCGGTGCGCCGGACCCAGCCGGAGGCCAGCTACAGGCCCTGGCGC

			GGGACAACACGCAACTGCTCATCAACCAGCTGGCAGCTGCCAACGGAGCCGTGGAAG AGCGATAGTGGCGGGCTGCCGGAGCCCACACGCCTGCCGAGAGAAGCCTCTGC CCGACCGCGGCCACTTACACGCTGGCAGCAGTCGCCGCCATCGCCAGGGCATCCGCCCC AGAAGAAGACCAACCTGGTGTGGACGAGGTGAGTGGCCAGTGGCGGGCTGGGCT ACCAGCGCCCGGGACGACACCAAAGAATGGCTGATTGAGGTGCCCGCAATGCCGACC CCTGGAGGACCAAGTCGCAAGCGGATTCAAGCCAAGAAGGAAGGGTGGCAAGAACG AGCTGAACCGGCTCGTAACCTGGCCCGCGCACAAGATGCAGCTGCCAGCGCGCC GCTTGACCCCTACCGGACACCAGAGTAAGGAGGAGCTGGCCGCATGCAAGTGGCA AGGTCTCCACCGCCTCTGTGGCGCTTCAGGAGGCCCTCCCCAAGGAGAAGGTGCCCC GGGGCTCCGGCAAGAAAAGGAAGTTCAACCCCTTTCGGGACTTGAGCCGAGAAAA AGAACAGTTGAGCTGCTCGTGTATGAAACAGCAAGAACGCTCAGCTGGATGTGACTA GGGCCACCAATAAGCAGATGAGGGAGGAGCAGGAGGAGGCCAAGAGGAGGAAAA TGAGCCAGAAGGGCAAGAGAAGGGAGGCCAGGGGGCTGGGAGGCAAGAGGAAAGGGG GCCCGCCCAGCCAGGGAGGAGGAAAGGGGGCTGGGAGGCAAGATGAATTCTGGC CGCCTGGCTTGGTGGCAAGAGAAAAGGAGGACAGGCCAGGAGGAAAGAGGAGGAAGT AATAGTTCTAACTGTCGGACCCGTCTGTAACCAAGGACTATGAAATACTAAATGTTAAG TTCTAGGCAATTATACGGGACTCAGAAGGACCTGGCGCTGCCCTATTGAGTTAAAG GGACAGGATTGCCCTCCGTCAAGAAAGTATGTAAGTGGACTGCACAAATTATGTT TTTCCCACAACCGAGACTTGGAGATTAAGAACTTATTGAGGATTAAGAATTAGGGAA ATAATTGGTGGAAACCGGGATGAGTTCTATTCTAAACAGCCTTTTTTTCTTTTA ATGTTGGATATACGGCGAGGTAGAGTTGGCCATATTGAGACTTAGATTGACGTATAT GTTCTGCATTATTTACAACAAGTTGTATCAGAGCGGGAGTCGGGGAGGGAAA GAAAACAAACAGTTCAGAATTGAATAGGCAAGTGACTIONTTAAAGATTAAGTAATAAA GATGTTATCTAGTG
S000116	F48	175	GGGGGCAGAGGGAGCGAGCGGGGCCCTAGGGTCAAGAGGCCGGCGAGCAGATTG CGCTCGGGCGCTCTGGGAAGGGAGTCCGGAGCCAACAGGGGGCTCGCCCTCTGGCCCA GCCCTTCCGGAGCCAACAGGGGACTTCGCCCTCTGGCCAGGCCCTCCCGCTGATCCCCCAG TCAGCGGTCGCAAGCCTGCCATCCACGAAACTTGGCCATACTGCGGGCGTACACT TTGCACTTGAACTTACAACACCCGAGCAAGGAGCGACTCTCCGACGCCGGTCTCTGGAAAGGCTGTC TCTGCCATTGGGACACTTCCCCGCCGCTGCCAGGACCCGGTTCTCTGGAAAGGCTGTC CTTGAAGCTCCTTAGACGCTGGAGTTTCTGGGAAGTGGGAAAGCAGCCTCCCGCAG ATGCCCTCAACGTTAGCTTACCAACAGGAACATGACCTGACTACGACTCGGTGAG CCGTATTCTACTGCCACGAGGAGGAGAACTTCTACCAAGCAGCAGCAGAGCGAGCTG CAGCCCCCGGCCAGCGAGGATATCTGGAAAGAAATTGAGCTGCTGCCACCCGCC CTGCCCCCTAGCCGCGCTCCGGCTCTGCTGCCCTCCTACGGTGCCTCACACCCCTC TCCCTCGGGAGACAACGACGGCGGTGGCGGGAGCTCTCCACGGCCGACCAGCTGGAG ATGGTGACCGAGCTGCTGGAGGAGACATGGTAACCAAGAGTTCATCTGCACCCGGAC GACGAGACCTTCATAAAAACATCATCATCAGGACTGTATGTGGAGCGGCTCTGGCC GCCGCCAAGCTCGTCTAGAGAAGCTGGCCTCTACAGGCTCGCGCAAAGACAGCGC AGCCCGAACCCGCCGCGCCACAGCGCTGCTCCACCTCCAGCTGTACCTGCAGGAT CTGAGCGCCGCCGCTCAGAGTCATCGACCCCTCGTGGTCTCCCTACCCCTCAAC GACAGCAGCTGCCCAAGTCTGCGCTCGCAAGACTCCAGGCCCTCTCCGCTCG GATTCTCTGCTCTCCCTGACGGAGTCCTCCCGCAGGGCAGCCCGAGCCCTGGTGC CATGAGGAGACACCGCCCACCAACAGCAGCAGCTGAGGAGGAACAAGAAGATGAGGAA GAAATCGATGTTGTTCTGTTGAAAAGAGGCAGGCTCTGGCAAAAGGTCAAGAGTCTGGA TCACCTCTGCTGGAGGCCACAGCAAACCTCTCACAGCCCAGTGGCTCAAGAGGTGC CACGTCTCCACACATCAGCACAACAGCAGCAGCGCCCTCCACTCGGAAGGACTATCCT GCTGCCAAGAGGTCAAGTGGACAGTGTCAAGACTCTGAGACAGATCAGAACACCGA AAATGCACCAAGCCCCAGGTCTCGGACACCGAGGAGAATGTCAAGAGGCGAACACACAAC

			GTCTGGAGCGCCAGAGGAGGAACGAGCTAAACGGAGCTTTTGCCTCGTGACCAG ATCCCGAGTTGAAAACAATGAAAAGGCCCAAGGTAGTTATCCTAAAAAGCCACA GCATACATCCTGTCCGTCCAAGCAGAGGAGCAAAGCTATTCTGAAGAGGACTTGTG CGGAAACGACGAGAACAGTTGAAACACAAACTGAAACAGCTACGGAACCTGTGCGTAA GGAAAAGTAAGGAAAACGATTCTTCAACAGAAATGTCCTGAGCAATCACCTATGAAC TGTTCAAATGCATGATCAAATGCAACCTCACAACCTGGCTGAGTCTTGAGACTGAAAG ATTTAGCCATAATGTAACAGTTGCTCAAATTGGACTTGGGATAAAAGAACCTTTATGC TTACCATCTTTTTCTTAAACAGATTGTATTAAAGAATTGTTAAAAAATTTT AAGATTACACAATGTTCTGTAAATATTGCCATTAAATGTAATAACTTAATAAAA ACGTTATAGCAGTTACACAGAATTCAATCCTAGTATATAGTACCTAGTATTAGGTA CTATAACCTAATTTTTATTAAAGTACATTGCTTTAAAGTTGATT
S000118	F49	176	GGGGGCAGAGGGAGCGAGCGGGCGGCCCTAGGGTCAAGAGGCCGGCGAGCAGAGTTG CGCTGCGGGCGTCTGGAGGGAGTTCCGGAGCCAACAGGGGCTTCGCCCTGGCCCA GCCCTCCGGAGCCAACAGGGGACTTCGCCCTGGCCCAGCCCTCCCGCTGATCCCCAG TCAGCGGTCCGCAAGCCTTGCACGAAACTTGGCCATACTGCCGAGCAGGGAGACTAT TTGCACTTGAACTTACAACACCCGAGCAAGGACGCGACTCTCCGACGCCGGAGACTAT TCTGCCCATTTGGGACACTTCCCGCCGCTGCCAGGACCCGGTCTCTGGAAAGGCTGTC CTTGAAGCTCCCTAGACGCTGGAGTTTCTGGGAAGTGGGAAAGCAGCCTCCCGCAGC ATGCCCTCAACGTTAGCTTCAACACAGGAACATGACCTCGACTACGACTCGTGCAG CCGTTACTGCGACGGAGGAGGAACCTTACCAAGCAGCAGCAGCAGGAGCTG CAGCCCCCGCGCCAGCGAGGATATCTGGAAAGAATTGAGCTGCTGCCACCCGCC CTGCCCCTAGCCGCCGCTCCGGCTCTGCTGCCCTCACGTTGCCGTACACCCCTC TCCCTCGGGAGACAACGACGGGGCTGGCAGGAGCTTCTCCACGCCGACAGCTGGAG ATGGTGACCGAGCTGGAGGAGACATGGTAACCAGAGTTCTGCGACCCGGAC GACAGACCTTCAAAACATCATCCAGGACTGTATGTGGAGCGGCTTCGGCC GCCCCAAGCTCGTCTCAGAGAACGCTGGCCTTACCAAGGCTGCCGCAAAGACAGCGC AGCCCACCCCGCCGCGCCAGCGCTGCTCCACCTCCAGCTGTACCTGCAGGAT CTGAGCGCCGCCGCTCAGAGTGCATGCCACCCCTGGTGTCTCCCTACCCCTCAAC GACAGCAGCTGCCCAAGCTCTGCCCTCGCAAGACTCCAGCGCCTCTCCGCTCG GATTCTCTGCTCTCCCGACGGAGTCTCCCGCAGGGCAGCCCCGAGCCCTGGTGC CATGAGGAGACACCGCCACCAACCGCAGCAGCAGCTTGAGGAGGAACAAGAAGATGAGGAA GAAATGATGTTCTGAGGGCTCTGGAAAAGAGGCAGGCTCTGGAAAAGGTCAGAGTCTGG TCACCTCTGCTGGAGGCCACAGCAAACCTCTCACAGCCCCTGGCTCAAGAGGTGC CACGTCTCCACACATCAGCACAACTACGCAGCGCCTCCCTCCACTCGGAAGGACTATC GCTGCCAAGAGGGCTAAGTGGACAGTGTCAAGAGTCTGAGACAGATCAGCAACAACGA AAATGCACCAAGCCCCAGGTCTCGGACACCGAGGAGGAATGTCAAGAGGCGAACACACAAC GTCTGGAGGCCAGAGGAGGAACGAGCTAAACGGAGCTTTTGCCTCGTGACCAG ATCCCGAGTTGAAAACAATGAAAAGGCCCAAGGTAGTTATCCTAAAAAGCCACA GCATACATCCTGTCCGTCCAAGCAGAGGAGCAAAGCTATTCTGAAGAGGACTTGTG CGGAAACGACGAGAACAGTTGAAACACAAACTGAAACAGCTACGGAACCTGTGCGTAA GGAAAAGTAAGGAAAACGATTCTTCAACAGAAATGTCCTGAGCAATCACCTATGAAC TGTTCAAATGCATGATCAAATGCAACCTCACAACCTGGCTGAGTCTTGAGACTGAAAG ATTTAGCCATAATGTAACAGTTGCTCAAATTGGACTTGGGATAAAAGAACCTTTATGC TTACCATCTTTTTCTTAAACAGATTGTATTAAAGAATTGTTAAAAAATTTT AAGATTACACAATGTTCTGTAAATATTGCCATTAAATGTAATAACTTAATAAAA ACGTTATAGCAGTTACACAGAATTCAATCCTAGTATATAGTACCTAGTATTAGGTA CTATAACCTAATTTTTATTAAAGTACATTGCTTTAAAGTTGATT
S000121	F50	177	GGGGGCAGAGGGAGCGAGCGGGCGGCCCTAGGGTCAAGAGGCCGGCGAGCAGAGTTG CGCTGCGGGCGTCTGGAGGGAGTTCCGGAGCCAACAGGGGCTTCGCCCTGGCCCA

			GCCCTTCCGGAGCCAACAGGGACTTCGCCTCTGGCCCAGCCCTCCGCTGATCCCCAG TCAGCGGTCCGCAAGCCTGCCGATCCACGAAACTTGCCTACTGCGGGCGTACACT TTGCACTTGAACCTACAACACCCGAGCAAGGACGCGACTCTCCCGACGCCGGAGACTAT TCTGCCCATTTGGGACACTTCCCGCCGCTGCCAGGACCCGGTCTCTGGAAGGCTGTC CTTGAAGCTCCTAGACGCTGGAGTTTTCGGGAAGTGGAAAGCAGCCTCCGCGACG ATGCCCTCAACGTTAGCTCACCAACAGGAACATGACCTGACTACGACTCGGTGAG CCGTATTCTACTGCGACGAGGAGGAGAACCTTACCAAGCAGCAGCAGAGCAGCTG CAGCCCCCGCGCCAGCGAGGATATGGAAGAAATTGAGCTGCTGCCACCCGGCC CTGCCCCTAGCCGCCCTCCGGCTCTGCTGCCCTCTACGTTGGTACACCCTTC TCCCTTCGGGAGACAACGACGGCGTGGCGGGAGCTCTCCACGGCCGACCAAGCTGGAG ATGGTGACCGAGCTGCTGGAGGAGACATGGTAACCAAGAGTTATCTGCAACCCGGAC GACGAGACCTTCATCAAAACATCATCATCCAGGACTGTATGTGGAGCGGCTTCGGCC GCCGCCAAGCTCGTCTCAGAGAAGCTGGCCTCTACCAAGGCTGCGCGCAAAGACAGCGGC AGCCCGAACCCGCCGGCCACAGCGTCTGCTCACCTCCAGCTTGTACCTGCAGGAT CTGAGCGCCGCCCTCAGAGTCATCGACCCCTGGTGGTCTCCCTACCCCTCAAC GACAGCAGCTGCCCAAGTCCTGCGCTCGCAAGACTCCAGCGCCTCTCCGCTCG GATTCTCTGCTCTCCTGACGGAGTCCTCCCGCAGGGCAGCCCCGAGCCCCGGTCTC CATGAGGAGACACCGCCCACCACAGCAGCAGCTGAGGAGGAACAAGAAGATGAGGAA GAAATCGATGTTGTTCTGTGGAAAAGAGGGCAGGCTCTGGCAAAAGGTCAAGAGTCTGG TCACCTTCTGCTGGAGGCCACAGCAAACCTCTCACAGCCCCTGGCTCAAGAGGTGC CACGTCTCCACACATCAGCACAACTACGCAGCGCCTCCACTCGGAAGGACTATCCT GCTGCCAAGAGGGTCAAGTTGGACAGTGTCAAGACTCTGAGACAGATCAGCAACAAACCGA AAATGCACCAAGCCCCAGGTCTCGACACCGAGGAGAATGTCAAGAGGGCAACACACAAC GTCTGGAGCGCCAGAGGAGGAACGAGCTAAACGGAGCTTTTGCCCTGCGTACCGAG ATCCCGGAGTTGAAAACATGAAAAGGCCCCAAGGTAGTTATCCTAAAAAGCCACA GCATACATCCTGTCGCTCAAGCAGAGGAGCAAAGCTCATTCTGAAGAGGACTTGTG CGGAAACGACGAGAACAGTTGAAACACAAACTGAACAGCTACGGAACCTTGCGTAA GGAAAAGTAAGGAAAACGATTCTTAACAGATTGTATTTAAGAATTGTTAAAAAATTT TGTTTCAATGCATGATCAAATGCAACCTCACAAACCTGGCTGAGTCTTGAGACTGAAAG ATTAGCCATAATGTAACACTGCCTCAAATTGGACTTGGCATAAAAGAAACTTTTATGC TTACCATTTTTTTCTTAACAGATTGTATTTAAGAATTGTTAAAAAATTT AAGATTACACAATGTTCTGTAAATATTGCCATTAAATGTAATAACTTTAATAAAA ACGTTTATGAGCAGTTACACAGAATTCAATCCTAGTATATAGTACCTAGTATTAGGTA CTATAAACCTAATTTTTATTAAAGTACATTGCTTTAAAGTTGATT
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A Pik3r1 nucleic acid sequence of the invention is depicted in Table 4 as SEQ ID NO. 178. The nucleic acid sequence shown is from mouse. SEQ ID NO: 179 (Table 5) depicts the amino acid sequence encoded by SEQ ID NO: 178. SEQ ID NO: 178 and SEQ ID NO: 179 are from mouse.

TABLE 4

SEQ. ID NO.	MOUSE SEQUENCE
178	GGCACGAGCC GAGTTGGAGG AAGCAGCGGC AGCGGCAGGC GCAGCGGTAG CGGTGAGGAC GGCTGTGCAG CCAAGGAACC GGGACAGCGA AGCGACGGCA GGTCGAGCT GGATCGCAGG AGCCTGGGAG CTGGGAGCTT CAGAGGCCGC TGAAGCCCAG GCTGGCAGA GGAAGGAAGC GAGCCGACCC GGAGGTTGAAG CTGAGAGTGG AGCGTGGCAG TAAAATCAGA CGACAGATGG ACAGTGTGAC AGGAACGTCA GAGAGGATTG GGCTCGCTG CGAGAGTCAG CCTGGAGTCA AGGTGTTGAC AAGTTGCTGA GAAGGACACG TGGGAGGACG GTGGCGCGCG GAGGGAGAG CCTGCTTCA GTACACCCGT TGATGGAGGA CAGATGGACA GCAGCCGGAC GGCCAGTCAC CTCTCTTAA CCTTGGATA GTGGCCTTT GTGCTCTGCT GGACACCTGT TGGGGATTT AGCCCATTCT CTGAACTCAC TTTCTCTTAA AACGTAACACT CGGACGGCAG TGTGCGAGCC AGCTCCTCTG TGGCAGGGCA CTAGAGCTGC AGACATGAGT GCAGAGGGCT ACCAGTACAG AGCACTGTAC GACTACAAGA AGGAGCGAGA GGAAGACATT GACCTACACC TGGGGGACAT ACTGACTGT AATAAAGGCT CCTTAGTGGC ACTTGATTGATTC AGTGATGGCC AGGAAGCCCG GCCTGAAGAT ATTGGCTGGT TAAATGGCTA CAATGAAACCC ACTGGGGAGA GGGGAGACTT TCCAGGAAC TACGTTGAAT ACATTGGAAG GAAAAGAATT TCACCCCCCTA CTCCAAGCC TCGGCCCCCT CGACCGCTTC CTGTTGCTCC GGGTTCTTCA AAAACTGAAG CTGACACCGA GCAGCAAGCG TTGCCCCCTC CTGACCTGGC CGAGCAGTTT GCCCCTCTG ATTTGCCCC GCCTCTCTT ATAAGCTCC TGGAAAGCCAT TGAGAAGAAA GGACTGGAAT GTTCGACTCT ATACAGAACAA CAAAGCTCCA GCAACCCCTGC AGAATTACGA CAGCTCTTG ATTGTGATGC CGCGTCAGTG GACTGGAGA TGATCGACGT ACACGTCTTA GCAGATGCTT TCAAACGCTA TCTCGCCGAC TTACCAAATC CTGTCATTCC TGTAGCTGTT TACAATGAGA TGATGTCTT AGCCCAAGAA CTACAGAGCC CTGAAGACTG CATCCAGCTG TTGAAGAACG TCATTAGATT GCCTAATATA CCTCATCAGT GTTGGCTTAC GCTTCAGTAT TTGCTCAAGC ATTTTCAA GCTCTCTCAA GCCTCCAGCA AAAACCTTTT GAATGCAAGA GTCTCTCTG AGATTTCAAG CCCCGTGTCTT TTCAGATTTC CAGCCGCCAG CTCTGATAAT ACTGAACACC TCATAAAAGC GATAGAGATT TTAATCTCAA CGGAATGGAA TGAGAGACAG CCAGCACCAAG CACTGCCCC CAAACCCACCC AAGCCCACTA CTGTAGCCAA CAACAGCATG AACACAATA TGCTCTTGCA GGATGCTGAA TGGTACTGGG GAGACATCTC AAGGGAGAGA GTGAATGAAA AACTCCGAGA CACTGCTGAT GGGACCTTTT TGTTACGAGA CGCATCTACT AAAATGCACG GCGATTACAC TCTTACACCT AGGAAAGGAG GAAATAACAA ATTAATCAAA ATCTTCACC GTGATGGAAA ATATGGCTTC TCTGATCCAT TAACCTCTCAA CTCTGTGGTT GAGTTAAATA ACCACTACCG GAATGAGTCT TTAGCTCAGT ACAACCCCCA GCTGGATGTG AAGTTGCTCT ACCCAGTGTG CAAATACCAG CAGGATCAAG TTGTCAAAGA AGATAATATT GAAGCTGTAG GGAAAAAAATT ACATGAATAT AATACTCAAT TTCAAGAAAA AAGTCGGAA TATGATAGAT TATATGAGGA GTACACCCGT ACTTCCCAGG AAATCCAAAT GAAAAGAACG GCTATCGAAG CATTAAATGA AACCATAAAAA ATATTGAAAG AACAAATGCCA AACCCAGGAG CGGTACAGCA AAGAATACAT AGAGAAGTT AAACGCGAAG GCAACGAGAA AGAAATCAA AGGATTATGC ATAACCATGA TAAGCTGAAG TCGCGTATCA GTGAGATCAT TGACAGTAGG AGGAGGTTGG AAGAAGACTT

SEQ. ID NO.	MOUSE SEQUENCE
	GAAGAAGCAG GCAGCTGAGT ACCGAGAGAT CGACAAACGC ATGAACAGTA TTAAGCCGGA CCTCATCCAG TTGAGAAAAGA CAAGAGACCA ATACTGTGATG TGGCTGACGC AGAAAGGTGT GCGGCAGAAG AAGCTAACG AGTGGCTGGG GAATGAAAAT ACCGAAGATC AATACTCCCT GGTAGAAGAT GATGAGGATT TGCCCCACCA TGACGAGAAG ACCTGGAATG TCGGGAGCAG CAACCGAAAC AAAGCGGAGA ACCTATTGCG AGGGAAGCGA GACGGCACTT TCCTTGCCG GGAGAGCAGT AAGCAGGGCT GCTATGCCTG CTCCGTAGTG GTAGACGGCG AAGTCAAGCA TTGCGTCATT ACAAGAGCTG CCACCGGCTA TGGCTTGCC GAGCCCTACA ACCTGTACAG CTCCCTGAAG GAGCTGGTGC TACATTATCA ACACACCTCC CTCGTGCAGC ACAATGACTC CCTCAATGTC AACTAGCAT ACCCAGTATA TGCACAACAG AGGCATGAA GCGCTGCCCT CGGATCCAGT TCCTCACCTT CAAGCCACCC AAGGCCTCTG AGAAGCAAAG GGCTCCTCTC CAGCCCCGACC TGTGAACCTGA GCTGCAGAAA TGAAGCCGGC TGCTGCACA TGGGACTAGA GCTTCTTGG ACAAAAAGAA GTCGGGAAAG ACACCGAGCC TCGGACTGTT GGATGACCAAG ACGTTTCTAA CCTTATCCTC TTCTTCTTCTT TCTTCTTCTT TCTTCTTCTT TCTTCTTCTT TCTTCTTCTT TTCTAATTT AAAGCCACAA CACACACCA ACACACAGAG AGAAAGAAAT GCAAAAATCT CTCCGTGCAG GGACAAAGAG GCCTTAACC ATGGTGCTT TTAACGCTT CTGAAGCTT ACCAAGCTACA AGTTGGACT TTGGAGACCA GAAGGTAGAC AGGGCCGAAG AGCCTGCGCC TGGGGCCGCT TGGTCCAGCC TGGTGTAGCC TGGGTGTCGC TGGGTGTGGT GAACCCAGAC ACATCACACT GTGGATTATT TCCTTTTAA AAGAGCGAAT GATATGTATC AGAGAGCCGC GTCTGCTCAC GCAGGACACT TTGAGAGAAC ATTGATGCAG TCTGTTCGGA GGAAAAATGA AACACCAAGAA AACGTTTTG TTTAAACTTA TCAAGTCAGC AACCAACAAAC CCACCAACAG AAAAAAAAAA AAAA

TABLE 5

MOUSE SEQUENCE	
179	MSAEQYQYRALYDYKKEREEDIDLHLGDLTVNKGSVALGFSD GQEARPEDIGWLNGYNETTGERGDFPGTYVEYIGRKIRSPPTPKPRPPRPLPVAPGSS KTEADTEQQALPLPDLAEQFAPPDVAPPILLKLEAIIEKKGLECSTLYRTQSSNPAE LRQLDCDAASVDEMDVHVLADAFKRYLADLPNPVIPVAVVNEMMSLAQELQSPED CIQLKKLIRLPNIPHQCWLTLQYLLKHFFKLSQASSKNLLNARVLSEIFSPVLFRFP AASSDNTHELIKAIIEILISTEWNERQPAPALPPKPPKPTTVANNNSMNNNMSLQDAEWY WGDISREEVNEKLRDTADGTFVLVDASTKMHGDYLTTPRKGGNNKLIKIFHRDGKYGF SDPLTFNSVVELINHYRNESLAQYNPKLDVKLLYPVSKYQQDQVVKEDNIEAVGKKLH EYNTQFQEKSREYDRLYEEYTRTSQEIQMKRTAIEAFNETIKIFEQCTQERYSKEY IEKFKREGNEKEIQRIMHNHDKLKSRISEIIDSRRRLLEEDLKQAAEYREIDKRMNSI KPDLIQLRKTRDQYLMWLTQKGVRQKKLNEWLGNENTEDQYSLVEDDEDLPHHDEKTW NVGSSRNKAENLLRGKRDGTFLVRESSKQGCYACSVVVDGEVKHCVINKTATGYGFA EPYNLYSSLKELVLHYQHTSLVQHNDSLNVTLAYPVYAQQR

Also suitable for use in the present invention is the sequence provided in Genbank Accession No. 5 U50413 and AAC52847.

Table 6 (SEQ ID NO: 180) depicts the nucleotide sequence of human Pik3r1. Table 7 (SEQ ID NO:181) depicts the amino acid sequence of human Pik3r1.

TABLE 6

HUMAN	
SEQ ID#	SEQUENCE
5 180	TACAACCAGG CTCAACTGTT GCATGGTAGC AGATTTGCAA ACATGAGTGC TGAGGGGTAC CAGTACAGAG CGCTGTATGA TTATAAAAAG GAAAGAGAAG AAGATATTGA CTTGCACTTG GGTACATAT TGACTGTGAA TAAAGGGTCC TTAGTAGCTC TTGGATTCAAG TGATGGACAG GAAGCCAGGC CTGAAGAACAT TGGCTGGTTA AATGGCTATA ATGAAACCAC AGGGGAAAGG GGGGACTTTG CGGGAACTTA CGTAGAACAT ATTGGAAAGGA AAAAACCTC GCCTCCCACA CCAAGCCCC GCCCACCTCG GCCTCTTCCT GTTGCACCAAG GTTCTCGAA AACTGAAGCA GATGTTAAC AACAAAGCTTT GACTCTCCCG GATCTTGCAAG AGCAGTTGC CCCTCCTGAC ATTGCCCGC CTCTTCTTCAAGCTCGT GAAGCCATTG AAAAGAAAGG TCTGGAATGT TCAACTCTAT ACAGAACACA GAGCTCCAGC AACCTGGCAAG ATTACGACA GCTTCTTGAT TGTGATACAC CCTCCGTGGA CTTGGAAATG ATCGATGTGC ACGTTTGGC TGACGCTTCA AAACGCTATC TCCTGGACTT ACCAAATCCT GTCATCCAG CAGCCGTTA CAGTGAATG ATTTCTTAG CTCCAGAACAT ACAAAAGCTCC GAAGAACATATA TTCAGCTATT GAAGAACGTT ATTAGGTCGC CTAGCATACC TCATCACTAT TGGCTTACGC TTCACTTATTT GTAAAACAT TTCTCAAGC TCTCTCAAAC CTCCAGCAAA AATCTGTTGA ATGCAAGAGT ACTCTCTGAA ATTTCTAGCC CTATGCTTT CAGATTCTCA GCAGCCAGCT CTGATAATAC TGAAAACCTC ATAAAAGTTA TAGAAATTTT AATCTCAACT GAATGGAATG AACGACAGCC TGCACCCAGCA CTGCCTCTA AACCACCAAA ACCTACTACT GTAGCCAACA ACGGTATGAA TAACAATATG TCCTTACAAA ATGCTGAATG GTACTGGGGA GATATCTCGA GGGAAAGAAGT GAATGAAAAA CTTCGAGATA CAGCAGACGG GACCTTTTG GTACGAGATG CGTCTACTAA AATGCATGGT GATTATACTC TTACACTAAG GAAAGGGGGA AATAACAAAT TAATCAAAAT ATTTCTATCGA GATGGAAAT ATGGCTTCTC TGACCCATTACCTCAGTT CTGTGGTTGA ATTAATAAAC CACTACCGGA ATGAATCTCT AGCTCACTAT AATCCAAAT TGGATGTGAA ATTACTTAT CCAGTATCCA AATACCAACA GGATCAAGTT GTCAAAGAAG ATAATATTGA AGCTGTAGGG AAAAAAATTAC ATGAATATAA CACTCAGTTT CAAGAAAAAA GTCGAGAATA TGATAGATTA TATGAAGAACAT ACCACCGCAC ATCCACAGGA ATCCAAATGA AAAGGACAGC TATTGAAGCA TTAATGAAA CCATAAAAAT ATTTGAAGAA CAGTGCAGA CCCAAGAGCG GTACAGCAAA GAATACATAG AAAAGTTAA ACGTGAAGGC AATGAGAACG AAATACAAAG GATTATGCAT AATTATGATA AGTTGAAGTC TCGAACATCACT GAAATTATTG ACAGTAGAACG AAGATTGGAA GAAGACTTGA AGAAGCAGGC AGCTGAGTAT CGAGAACATTG ACAAACGTAT GAACAGCATT AAACCAAGACC TTATCCAGCT GAGAAAGACG AGAGACCAAT ACTTGATGTG GTTACTCAA AAAGGTGTTG GGCAAAAGAA GTTGAACGAG TGTTGGGCA ATGAAAACAC TGAAGACCAA TATTCACTGG TGGAAAGATGA TGAAGATTG CCCCACATCATG ATGAGAACAC ATGGAATGTT GGAAGCAGCA ACCGAAACAA AGCTGAAAAC CTGTTGCGAG GGAAGCGAGA TGGCACTTTT CTTGTCCGGG AGAGCAGTAA ACAGGGCTGC TATGCCCTGCT CTGTAGTGGT GGACGGCGAA GTAAAGCATT GTGTCACTAA CAAACACAGCA ACTGGCTATG GCTTGGCGA GCCCTATAAC TTGTACAGCT CTCTGAAAGA ACTGGTGCTA CATTACCAAC ACACCTCCCT TGTGCAGCAC AACGACTCCC TCAATGTCAAC ACTAGCCTAC CCAGTATATG CACAGCAGAG GCGATGAAGC GCTTACTCTT TGATCCCTCT CCTGAAGTTC AGCCACCCCTG AGGCCTCTGG AAAGCAAAGG GCTCCTCTCC AGTCTGATCT GTGAATTGAG CTGCAGAAC GAAGCCATCT TTCTTGGAT GGGACTAGAG CTTTCTTCA CAAAAAAGAA GTAGGGGAAG ACATGCAGCC TAAGGCTGTA TGATGACCAC ACGTTCTAA GCTGGAGTGC TTATCCCTTC TTTTCTTTT TTTCTTGGT TTAATTAAA GCCACAACCA CATAAACAC AAAGAGAAAA AGAAATGCAA AAATCTCTGC GTGCAGGGAC AAAGAGGGCCT TTAACCATGG TGCTTGTAA TGCTTCTGA AGCTTACCA

HUMAN	
SEQ ID#	SEQUENCE
	GCTGAAAGTT GGGACTCTGG AGAGCGGAGG AGAGAGAGGC AGAAGAACCC TGGCCTGAGA AGGTTGGTC CAGCCTGGTT TAGCCTGGAT GTTGCTGTGC ACGGTGGACC CAGACACATC GCACTGTGGA TTATTCATT TTGTAACAAA TGAACGATAT GTAGCAGAAA GGCACGTCCA CTCACAAGGG ACGCTTGGG AGAATGTCAG TTCATGTATG TTCAGAAGAA ATTCTGTCAT AGAAAGTGCC AGAAAGTGTGTT TAACTTGTCA AAAAACAAAA ACCCAGCAAC AGAAAAATGG AGTTTGGAAA ACAGGACTTA AAATGACATT CAGTATATAA AATATGTACA TAATATTGGA TGACTAACTA TCAAATAGAT GGATTGTAT CAATACAAA TAGCTTCTGT TTTGTTTGC TGAAGGCTAA ATTCACAGCG CTATGCAATT CTTAATTTC ATTAAAGTTGT TATTTCAGTT TTAAATGTAC CTTCAGAATA AGCTTCCCCA CCCCAGTTT TGTTGCTTGA AAATATTGTT GTCCCGGATT TTTGTTAATA TTCATTGGT TTATCCTTTT TAAAAAATAA ATGTACAGGA TGCCAGTAA AAAAATG GCTTCAGAAT TAAAACATG AAATATTAA CAGTTTTCT TGTACAGAGT ACTTGCTGTT AGCCCAAGGT TAAAAGTT ATAACAGATT TTTTTGGAC TGTTTGTG GGCACTGCCT GATAAGCTTC AAAGCTGCTT TATTCAATAA AAAAATG CGAATTCACT GG

TABLE 7

HUMAN SEQUENCE	
181	MSAEGYQYRA LYDYKKEREE DIDLHLGDIL TVNKGSVAL GFSDGQEARP EEIGWLNGYN ETTGERGDFP GTYVEYIGRK KISPPPKPR PPRPLPVAPG SSKTEADVEQ QALTLPDLAE QFAPPDIAPP LLIKLEVAIE KKGLECSTLY RTQSSNLAE LRQLLDCDTP SVDLEMIDVH VLADAFKRYL LDLPNPVIPA AVYSEMISLA PEVQSSEEEYI QLLKKLIRSP SIPHQYWLT QYLLKHFFKL SQTSSKNILLN ARVLSEIFSP MLFRSAASS DNTENLIKVI EILISTEWNE RQPAPALPPK PPKPTTVANN GMNNNMSLQN AEWYWGDISR EEVNEKLRDT ADGTFLVRDA STKMHGDYTL TLRKGNNKL IKIFHRDGKY GFSDPLTFSS VVELINHYRN ESLAQYNPKL DVKLILYPVSK YQQDQVVKED NIEAVGKKLH EYNTQFQEKS REYDRRLYEY TRTSQEIQMK RTAIEAFNET IKIFEEQCQT QERYSKEYIE KFKREGNEKE IQRIMHNYDK LKSRISEIID SRRRLEEDLK KQAAEYREID KRMNSIKPDL IQLRKTRDQY LMWLTQKGVR QKKLNEWLGN ENTEDQYSLV EDDDEDLPHHD EKTWNVGSSN RNKAENLLRG KRDGTFLVRE SSKQGCYACS VVVGEVKHC VINKTATGYG FAEPYNLYSS LKELVLHYQH TSLVQHNDSL NVTLAYPVYA QQRR

Also suitable for use in the present invention is the sequence provided in Genbank Accession No. 10 M61906 and A38748.

A GNAS nucleic acid sequence of the invention is depicted in Table 8 as SEQ ID NO. 182. The nucleic acid sequence shown is from mouse.

TABLE 8

TAG #	SEQ. ID NO.	SEQUENCE
S00056	182	GACGGTATGCAGTAGAAATAAGGTCTCAGCAGTCAGTGCAGAAAATCAAGCAAAGCCCC CTTAGGAGTTATTCATGTTGCCGCTTCGTGCAAATAGGGGAGGGGGCTTAAGGCTTACCG GAAGACCCCCCACCTAGCTCAGGTCTTGACTTCTGTCTGGTAAAGGCAAAGGAGATT TGGGGTGTAGTTGATGGCCATTAGGGTGGCTCGCAGACTAGAAAACCTGAAATGCACTTA AC

A contig assembled from the mouse EST database by the National Center for Biotechnology Information (NCBI) having homology with all or parts of the GNAS nucleic acid sequence of the invention is depicted 5 in Table 9 as SEQ ID NO. 183. SEQ ID NO. 184 represents the amino acid sequence of a protein encoded by SEQ ID NO. 183 and corresponds to mouse G protein Xl_{α^s} .

TABLE 9

MOUSE			
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
10 S00056	F12	183	GTTGAGCGCGAACAGCAGCCGAGATGGAAGGAAGCCCCTACCACCGCCACTGCGGTGGAAGGA AAAGTCCCCTCTCCGGAGAGAGGGGACGGATCTTCCACCCAGCCTGAAGCAATGGATGCC AAGCCAGCCCCCTGCTGCCAACAGCCCTCTACCGGATCTGATGCTGGAGCTCTACGGAT TCCCGCATGCTCACAGATAGCCAGAGCGATGCCGGAGAAAGACGGGACAGCCCCAGGAACG CCTTCAGATCTCCAGTCGGATCCTGAAGAACTCGAAGAAAGCCCCAGCTGTCCGCGCCGAT CCTGACGGAGGGGAGCCCCAGTCGCCCCAGCCACTCTGCCAGTCCGAGTCTGAAGGC AGCAGAGATCCAGCCGCCAGCCAGCCTCCGAGGCAGTCCCTGCCACCACGGGGAGTCT GCCTCCGGGGCAGCCCCCTGTCAcccAGGTGGAGCCGCAGCCGCGCAGTCTGCCACC CTGGCGGAGGCTGCCGCCGGCAGCCCCCTATCACCCCCAAGGAGCCCCTACCCGGGCA GTCCCCCTGCTAGAGCCCATCCGGCCGCTGGAGCAGTCCCTGGCGCCCCAGCAATGTCA GCCTCTGCTAGGGCAGCTGCCGCTAGGGCAGCCTATGCAGGTCCACTGGCTGGGGAGCC AGGTCACTCTCAGCTACTCCGCCGCTGGGCATCCCTTCTGCCGCCAGCAGCTGCG GCCCGGGCAGCCTCTGCCGCCAGTCGCTGCTGGCCGGTCAGCCTCTGCCGCC AGCAGGGCCCCTTAAGACCCCCCAGCCCCAGATCCAGGTTGCTGACCCGCTACTCCG CGGCTCCTCCGCCGCGACTGCCCTGACAAGTACGAGCAGGGCGAAGCTGCTGC AGGTACGAGGCATCGTCTGGCATCTCGAGATCGAGTCCTCCAGTGTAGTCGGAGAA GGGGCCACCGGCTGCTTCCAGTGGCTCTGCCGGAAACCCGCCCTGGCCTGCCCG AGCCACACGGTGGAGCAACCCAGTCCGCAACTTCTCACCGAGCCTCGGAAGCTGCG TTCGGTCTATCCGAGTGTACCCGATCACGATCCCTCAGCCCCGGAGGCCAAGGATCCT ATGGAGGAGAGGCAGCAACAGATGCCAAAGAAGCCATTGAGATGCGAGAGCAGAAGCGC GCAGATAAGAAACGCAGCAAGCTCATCGACAAGCAACTGGAGGAGGAGAAGATGGACTAC ATGTGTACACACCGCCTGCTGCTTAGGTGCTGGAGAGTCTGGCAAAGCACCATTGTG AAGCAGATGAGGATCCTGCATGTTAATGGTTAACGGAGATAGTGAGAAGGCCACTAA GTGCAGGACATCAAAACACCTGAAGGAGGCCATTGAAACCATTGTGGCCGCCATGAGC AACCTGGTGCCCCCTGTGGAGCTGCCAACCTGAGAACCAGTTCAAGGTGAGTGGACTACATT CTGAGCGTGTGAACTGCGAACATTGACTTCCCACCTGAATTCTATGAGCATGCCAAG GCTCTGTGGAGGAGTGGAGCTGCCGTGCTACGAGCGCTCCAATGAGTACCGCTG ATTGACTGTGCCAGTACTCCTGGACAAGATTGATGTGATCAAGCAGGCCACTACGTG CCAAGTGACCAGGACCTGCTCGCTGCCGTGCTGACCTCTGGAATCTTGAGACCAAG TTCCAGGTGGACAAAGTCAACTCCACATGTTGATGTGGCGGCCAGCGCGATGAGCGC

SAGRES TAG#	REF #	SEQ ID#	MOUSE
			SEQUENCE
			CGCAAGTGGATCCAGTGCTTCAATGATGTGACTGCCATCTCGTGGTGGCCAGCAGC AGCTACAAACATGGTCATTGGGAGGACAACCAAGACTAACCGCCTGCAGGAGGCTCTGAAC CTCTCAAGAGCATCTGGAACAACAGATGGCTGCGCACCATCTCTGATTCTCTCCCTC AACAAAGCAAGACCTGCTGCTGAGAAAAGTCCTCGCTGGCAAATCGAAGATTGAGGACTAC TTTCCAGAGTCGCTCGCTACACCACTCCTGAGGATGCGACTCCCGAGCCGGAGAGGAC CCACCGGTGACCCGGGCAAGTACTTCATTGGGATGAGTTCTGAGAATCAGCACTGCT AGTGGAGATGGGCGCCACTACTGCTACCCCTACTTACCTGCGCCGTGGACACTGAGAAC ATCCGCCGTCTTCAACGACTGCCGTGACATCATCCAGCGCATGCATCTCCGCCAAC GAGCTGCTCTAAGAAGGGAACACCCAATTAAATTCAAGCCTTAAGCACAATTAAAGA GTGAAACGTAATTGTACAAGCAGTGGTACCCACCATAAGGCATGATCAACACCGAAC CTTCCCTTTCCCCAGTGATTCTGAAAAACCCCTTCCCTCAGCTGCTTAGATGT TCCAAATTAGTAAGCTTAAGCGGCCCTACAGAAGAAAAAGAAAAAGGCCACAAAAG TTCCCTCTCACTTCAGTAAATAAAAGCAGCAACAGAAATAAGAAATAATGAA ATTCAAAATGAAATAATATTGTGTTGCAGCATTAAAAATCAATAAAATCAAAAAAT GAGCAAAAAAAAAAA
	184		MEGSPTTATAVEGVKVPSPERGDGSSTQPEAMDAKPAPAAQAVSTGSDAGAPTDASMLTDSQSD AGEDGTAPGTPSDLQSDPEELEEAPAVRADPDGAAPVAPATPAESESEGSRDPAEPAEAVP ATTAESASGAAPVTQVEPAAAASATLAEPARAAPITPKETTRAVPSARAHPAAGAVPGAPAM SASARAAAARAAYAGPLVWGARLSATPAARASLPARAAAARAASAARAVAAGRSASAAPSRA HLRPPSPEIQVADPPTPRPPRPTAWPDKYERGRSCCRYEASSGICEIESSSDESEEGATGCFQ WLLRRNRRPGLPRSHTVGSNPVRNFTRAFGSCFGLSECTRSRSLSPGKAKDPMEERRKQMRK EAIEMREQKRADKKRSKLIDKQLEEKMDYMCTHRLLLGAGESGKSTIVKQMRILHVNGFNGDS EKATKVQDIKNNLKEAIETIVAAMSNLVPPVELANPENQFRVDYILSVMNVPNFDFPPEFYEHAKAL WEDEGVRCACYERSNEYQLIDCAQYFLDKIDVIKQADYVPSDQDLLRCRVLTSGIFETKFQVDKVNF HMFDVGGQRDERRKWIQCFNDVTIAIFVVASSSYNMIREDNQTNRLQEALNLFSIWNNRWLRTI SVILFLNKQDLAELKVLAGSKIEDYFPEFARYTTPEDATPEPGEDPRVTRAKYFIRDEFLRISTASG DGRHYCYPHFTCAVDTENIRRVNDCRDIIQRMHLRQYELL

Also suitable for use in the present invention is Genbank Accession No. AF116268.

A contig assembled from the human EST database by the NCBI having homology with all or parts of the GNAS nucleic acid sequence of the invention is depicted in Table 10 as SEQ ID NO. 185. SEQ ID NO. 186 represents the amino acid sequence of a protein encoded by SEQ ID NO. 185 and corresponds to human G protein α_1 .

TABLE 10

SAGRES TAG#	REF #	SEQ ID#	HUMAN
			SEQUENCE
S000056	F37	185	ATGGAGACCGAACCGCCTCACAAACGAGCCCATCCCCGTCGAGAATGATGGCGAGGCTGT GGACCCCCAGAGGTCTCCAGACCCAACCTTCAGGTCTCAACCCGGCATTAGGGAAAGCT GGAGCCCAGCTACAGCCCACCTCCTGAGGAAGCAATGCCCTCGAGGCTGAACAG

HUMAN			
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
			CCCAGCTTGGGAGGGCTCTGGCCTACACTGGAGCAGCCTGGATTCCCCAGTGGGTCCAT GCAGGCCCTGCCAKGSTYSGSCCAGCACTCATGGAGCCGGAGCCTCAGTGGTGCAGA CCAGGCCCTGGGAGGATAACGCCCTCCACCAAGAAGAAAGCTATGCCCTTGAGTTGACCAG CCTGCCAGAGAGGCTGCAGTCACCTCTTACAGTCCAGACCTTGCTCCAGGAGGC CCAGGTGCTGCAGGGTCCCCGGAGCTCTCCAGGGAGGCCAAGCCTCAGGCCGCA AAGGCTGGCTCCAGAGGAGGCACAGCCCTCCCCCTGAGGAGACTATGCCATTGAGCTT GATGGAGAAGGATTGGGGACAGCAGCCACCCCCCGGGCTTCCGAGTTATCGCACAA GTCGACGGCAGCAGCCAGTTCGGCAGTCGCGCCTCGAGTGGGTCCGCTCACCTCC GCCGCAACGCGCCTCCCTCTGGGTCAGGCAGCATCGGAGCCATCCAAAGAGGCT GTCAGACCTCCTCTAACCTCACGGCAGCAGCCCTGGATGGAGATCTCCGACAGCCCC TTCGAGATTGGCAGCGCCCCCGCTGGGTCAGCAGACACTCCGTCAACATGGACAGCCCC CCAATCGCGCTTGACGGCCGCCATCAAGGTCTCCGGAGCCCCAGATAAGAGAGAGCGA GCAGAGAGACCCCCAGTTGAGGAGGAAGCAGCAGAGATGGAAGGAGCCGCTGATGCCGCG GAGGGAGGAAAAGTACCCCTCCGGGTACGGATCCCTGCCGCCGGGGCAGCCTCAGCG GATACCGCTGCCAGGGCAGCCCTGCAGCCCAAGCCGATCTGACTCCGGGCAACCCCA GAAGATCCCAGCTCCGGACAGCACCAGCCGATCTGACTCCGGGCACTCGCAGCCGAT CCCGACTCCGGGGCAGCCCTGCCGCCCCAGCCGATCCCGACTCCGGGCGCCCTGAC GCCCGAGCCGATCCGACTCCGGGCGCCCTGACGCCCCAGCCGATCCAGATGCCGG GCCGGCCCTGAGGCTCCGCCGCCCCCTGCCGCTGAGACCCGGCAGCCCATGCGCC CCAGCTGCCAGACGCGAGGGCTCCACTGCCCAAGCCGCTCTGCCACCCGGGAGCC CAAGTCCGCCGGCGGCCTCTGAGCCCTGCCCTCCGGGCAAGACGCAAGATCCATCTC AGACCCCCCAGCCCCAGATCCAGGCTGCCGATCCGCTACTCCGCGGCTACTCGCG TCTGCCCTGGGGCAAGTCCGAGAGCAGCCGCGGGCCGCGCTGTACTACGATGAAGGG GTGGCCAGCGAGCAGATGACTCCAGCGAGACGAGTCCGACGATGGGACCTCCGGATGC CTCCGCTGGTTTCAGCATGGCAAATGCCGCCGAAAGCCCCAGCCGAACTTACTC CGCAACTTCTCGTGCAAGCCTCGGGGCTGCTCGGTCATCTGAGAGTCCCCAGCCC AAAGCCTCGCGCTCTCAAGGTCAAGAAGGTACCCCTGGCGGAGAAGCGCAGACAGATG CGCAAAGAAGCCCTGGAGAAGCGGGCCCAGAGCGCGCAGAGAAGAAACGAGTAAGCTC ATCGACAAACAACCTCAGGACGAAAAGATGGCTACATGTGTACGCACCGCCTGCTGCTT CTAG
	186		MEISGPPFEIGSAPAGVDDTPVNMDSPPIALDGPPIKVSGAPDKRERAERPPVEEEAAEMEGAADA AEGGKVPSPGYGSAPAAGAASADTAARAAPAAPADPDSGATPEDPDSGATAPADPDSGAFAAADPDS GAAPAAAPADPDSGAAPDAPADPDSGAAPDAPADPDAGAAPEAPAAPAAAETRAAHVAPAAPDAG APTAPAASATRAAQVRRAASAAPASGARRKIHRLPPSPEIQAADPPTPRTRASAWRGKSESSRG RRVYYDEGVASSDDDSGDESDDGTSGCLRFQHRRNRRRKPQRNLLRNFLVQAFGGCFGRS ESPQPKASRSLSKVKKVPLAEKRRQMRKEALEKRAEKRSKLIDKQLQDEKMGYMCTHRLLL L

- 20 Table 11 demonstrates the nucleic acid sequence (SEQ ID NO: 187) and amino acid sequence (SEQ ID NO: 188) of NESP55 from mouse. SEQ ID NO: 188 represents the protein encoded by SEQ ID NO: 187.

TABLE 11

MOUSE			
SAGRES	REF	SEQ	SEQUENCE

TAG#	#	ID#	
	187		GAGAGGATCA GTGGAGGCAC CTCTCGGAGT CTTAGACTTC AGAGTCTGAG ACTTAGCGAG AGGAGCCTCG AGGAGACTCC TTCTCTCTTC TTTACCCATC CCTTTCTTTT ACTTACAGCC TCAAGCTGAG GCGCGGAGCT TTAGAAAGTT CGCAGTGTT TGAGTCCTT GCGCAGTGGG GCCACTCTCT GCAGAGCCAG AGGGTGAGTC GGCTTCTCGG TGAGCACCTA AGAGAATGGA TCGCAGGTCC CGGGCTCAGC AGTGGCGCC AGCTGCCAT AATTACAACG ACCTGTGCC GCCCATAGGC CGCCGGGCTG CCACCGCTCT CCTCTGGCTC TCCTGCTCCA TTGCTCTCCT CCGCGCCCTA GCCTTCTCCA ACGCCCGCAG CCAGCAGCGT GCTGCCCATC GCCGGAGCTT CCTTAACGCC CACCACCGCT CCGCTGCCGC TGCAAGCTGCC GCACAGGTAC TCCCTGAGTC CTCTGAATCT GAGTCTGATC ACGAGCACGA GGAGGTTGAG CCTGAGCTGG CCCGCCCCGA GTGCCTAGAG TACGATCAGG ACGACTACGA GACCGAGACC GATTCTGAGA CCGAGCCTGA GTCCGATATC GAATCCGAGA CGAAATCGA GACCGAGCCA GAGACCGAGC CAGAAACCGA GCCAGAGACC GAGCCAGAGG ACCAGCGCGG CCCCCGGGGT GCCACCTTCA ACCAGTCACT CACTCAGCGT CTGCACGCTC TGAAAGTTGCA GAGCGCCGAC GCCTCCCCGA GACGTGCGCA GCCCACCACT CAGGAGCCTG AGAGCGCAAG CGAGGGGGAG GAGCCCCAGC GAGGGCCCTT AGATCAGGAT CCTCGGGACC CCGAGGAGGA GCCAGAGGAG CGCAAGGAGG AAAACAGGCA GCCCCGCCGC TGCAAGACCA GGAGGCCAGC CCGCCGTCGC GACCAGTCCC CGGAGTCCCC TCCCAGAAAG GGGCCCATCC CCATCCGGCG TCACTAATGG GTGACTCCGT CCAGATTCTC CTTGTTTCA TGGATAAAGG TGCTGGAGAG TCTGGCAAAA GCACCATTTGT GAAGCAGATG AGGATCCTGC ATGTTAATGG GTTAAACCGGA G
	188		MDRRSRAQQWRRARHNYNDLCPPIGRRAATALLWLSCSIALLRA LASSNARAQQRAHRR SFLNAHHRSAAAAAAAQVLPESSESESDEHEHEEVEPELARPE CLEYDQDDYETETDSETEPESDIE SETEIETEPETEPETEPETEPEDERGPRGATFNQLSTQRLHALKLQSADASPRRAQPTTQEPESSA EGEEPQRGPLDQDPRDPEEEPEERKE ENRQPRRKTRRPARRRDQSPESPPRKGPIPIRRH

Table 12 demonstrates the nucleic acid sequence (SEQ ID NO: 189) and amino acid sequence (SEQ ID NO: 190) of NESP55 from human. SEQ ID NO: 190 represents the protein encoded by SEQ ID NO: 189.

TABLE 12

HUMAN			
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
5		189	CTCGCCTCAG TCTCCTCTGT CCTCTCCCAG GCAAGAGGAC CGGCGGAGGC ACCTCTCTCG AGTCTTAGGC TGCGGAATCT AAGACTCAGC GAGAGGAGCC CGGGAGGAGA CAGAACTTT CCCTTTTTC CCATCCCTTC TTCTTGCTCA GAGAGGCAAG CAAGGCGCGG AGCTTTAGAA AGTTCTTAAG TGGTCAGGAA GGTAGGTGCT TCCCTTTTC TCCTCACAAAG GAGGTGAGGC TGGGACCTCC GGGCCAGCTT CTCACCTCAT AGGGTGTACC TTCCCGGCT CCAGCAGCCA ATGTGCTTCG GAGCCGCTCT CTGCAGAGCC AGAGGGCAGG CGGCGTCTC GGTGTGTGCC TAAGAGGATG GATCGGAGGT CCCCCGGCTCA GCAGTGGCGC CGAGCTCGCC ATAATTACAA CGACCTGTGC CGGCCATAG GCCGCCGGGC AGCCACCGCG CTCCCTGGC TCTCCTGCTC CATCGCGCTC CTCCGCGCCC TTGCCACCTC CAACGCCCGT GCCCAGCAGC GCGCGGCTGC CCAACAGCGC CGGAGCTTCC TTAACGCCCA CCACCGCTCC GGCGCCAGG TATTCCCTGA GTCCCCCGAA TCGGAATCTG ACCACGAGCA CGAGGAGGCA GACCTTGAGC TGTCCTCCC CGAGTGCCTA GAGTACGAGG AAGAGTTCGA CTACGAGACC GAGAGCGAGA CCGAGTCCGA

HUMAN			
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
			AATCGAGTCC GAGACCGACT TCGAGACCGA GCCTGAGACC GCCCCCACCA CTGAGCCCGA GACCGAGCCT GAAGACGATC CGGGCCCGGT GGTGCCCAAG CACTCCACCT TCGGCCAGTC CCTCACCCAG CGTCTGCACG CTCTCAAGTT GCGAAGCCCC GACGCCCTCCC CAAGTCGCGC GCCGCCAGC ACTCAGGAGC CCCAGAGCCC CAGGGAAGGG GAGGAGCTCA AGCCCAGGA CAAAGATCCA AGGGACCCCG AAGAGTCGAA GGAGCCCAAG GAGGAGAAGC AGCGGCGTCG CTGCAAGCCA AAGAAGCCCA CCCGCCGTGA CGCGTCCCCG GAGTCCCCTT CCAAAAAGGG ACCCATCCCC ATCCGGCGTC ACTAATGGAG GACGCCGTCC AGATTCTCT TGTTTCATG GATTCAAGTG CTGGAGAACAT TGTTAAAGC ACCATTGTGA AGCAGATGAG GATCCTGCAT GTTAATGGGT TTAATGGAGA GGGCGGCGAA GAGGACCCGC AGGCTGCAAG GAGAACAGC GATGGCAGTG AGAAGGCAAC CAAAGTCAG GACATCAAAA ACAACCTGAA AGAGGCGATT GAAACCATTG TGGCCGCCAT GAGCAACCTG GTGCCCGCCG TGGAGCTGGC CAACCCCGAG AACCAAGTCA GAGTGGACTA CATCCTGAGT GTGATGAACG TGCGTACTT TGACTTCCCT CCCGAATTCT ATGAGCATGC CAAGGCTCTG TGGGAGGATG AAGGAGTGC G TGCGTCTAC GAACGCTCCA ACGAGTACCA GCTGATTGAC TGTGCCAGT ACTTCCTGGA CAAGATCGAC GTGATCAAGC AGGCTGACTA TGTGCCGAGC GATCAGGACC TGCTTGCCTG CCGTGTCCCTG ACTTCTGGAA TCTTGAGAC CAAGTCCAG GTGGACAAAG TCAACTTCCA CATGTTGAC GTGGGTGGCC AGCGCGATGA ACGCCGCAAG TGGATCCAGT GCTTCAACGA TGTGACTGCC ATCATCTCG TGGTGGCCAG CAGCAGCTAC AACATGGTCA TCCGGGAGGA CAACCAGACC AACCGCCTGC AGGAGGCTCT GAACCTCTTC AAGAGCATCT GGAACAACAG ATGGCTGCGC ACCATCTCTG TGATCCTGTT CCTCAACAAAG CAAGATCTGC TCGCTGAGAA AGTCCTTGCT GGGAAATCGA AGATTGAGGA CTACTTCCA GAATTGCTC GCTACACTAC TCCTGAGGAT GCTACTCCCG AGCCCGGAGA GGACCCACGC GTGACCCGGG CCAAGTACTT CATTGAGAT GAGTTCTGA GGATCAGCAC TGCCAGTGGA GATGGCGTC ACTACTGCTA CCCTCATTT ACCTGCGCTG TGGACACTGA GAACATCCGC CGTGTGTTCA ACGACTGCCG TGACATCATT CAGCGCATGC ACCTCGTCA GTACGAGCTG CTCTAAGAAG GGAACCCCCA ATTAAATTAA AAGCCTTAAG CACAATTAAAT TAAAAGTGA ACGTAATTGT ACAAGCAGTT AATCACCCAC CATAGGGCAT GATTAACAAA GCAACCTTTC CCTCCCCCG AGTGATTTG CGAAACCCCCC TTTCCCTTC AGCTTGCTTA GATGTTCCAA ATTTAGAAAG CTTAAGGCGG CCTACAGAAA AAGGAAAAAA GGCCACAAAA GTTCCCTCTC ACTTCAGTA AAAATAAAATA AAACAGCAGC AGCAAACAAA TAAAATGAAA TAAAAGAAAC AAATGAAATA AATATTGTGT TGTGAGCAT TAAAAAAAT CAAAATAAAA ATAAATGTG AGCAAAGAAA AAAAAA GAGAGGATCA GTGGAGGCAC CTCTCGGAGT CTTAGACTTC AGAGTCTGAG ACTTAGCGAG AGGAGCCTCG AGGAGACTCC TTCTCTCTTC TTACCCATC CCTTCTTT ACTTACAGCC TCAAGCTGAG GCGCGGAGCT TTAGAAAGTT CGCAGTGGTT TGAAGTCCTT GCGCAGTGGG GCCACTCTCT GCAGAGCCAG AGGGTGANTC GGCTTCTCG TGAGCACCTA AGAGAATGGA TCGAGGTCC CGGGCTCAGC AGTGGCGCCG AGCTGCCAT ATTACAACG ACCTGTGCC GCCCATAGGC CGCCGGGCTG CCACCGCTCT CCTCTGGCTC TCCTGCTCCA TTGCTCTCT CCGCGCCCTA GCCTCTTCCA ACAGCCCGCGC CCAGCAGCGT GCTGCCATC GCCGGAGCTT CCTTAACGCC CACCACCGCT CCGCTGCCGC TGAGCTGCC GCACAGGTAC TCCCTGAGTC CTCTGAATCT GAGTCTGATC ACGAGCACGA GGAGGTTGAG CCTGAGCTGG CCCGCCCCGA GTGCCTAGAG TACGATCAGG ACGACTACGA GACCGAGACC GATTCTGAGA CCGAGCCTGA GTCCGATATC GAATCCGAGA CCGAAATCGA GACCGAGCCA GAGACCGAGC CAGAAACCGA GCCAGAGACC GAGCCAGAGG ACGAGCGCGG CCCCCGGGGT GCCACCTTCA ACCAGTCACT CACTCAGCGT CTGCACGCTC TGAAGTTGCA GAGGCCGAC GCCTCCCCGA GACGTGCGCA GCCCACCCT CAGGAGCCTG AGAGCGCAAG CGAGGGGGAG

SAGRES TAG#	REF #	SEQ ID#	HUMAN
			SEQUENCE
			GAGCCCCAGC GAGGGCCCTT AGATCAGGAT CCTCGGGACC CCGAGGAGGA GCCAGAGGAG CGCAAGGAGG AAAACAGGCA GCCCCGCCGC TGCAAGACCA GGAGGCCAGC CCGCCGTCGC GACCAGTCCC CGGAGTCCCC TCCCAGAAAG GGGCCCATCC CCATCCGGCG TCACTAATGG GTGACTCCGT CCAGATTCTC CTTGTTTCA TGATAAAGG TGCTGGAGAG TCTGGCAAAA GCACCATTGT GAAGCAGATG AGGATCCTGC ATGTTAATGG GTTTAACCGGA G
	190		MDRRSRAQQWRRARHNYNDLCPPIGRRAATALLWLSCSIALLRA LATSNARAQQRAAAQQRSPFLNAHHRSGAQVFPESESDHEHEEADLESLPECLE YEEEFDYETESETESEIESETDFETEPETAPTTEPETEPEDDORGPVVKHSTFGQS LT QRLHALKLSPDASPSPRAPPSTQEPQSPREGEELKPEDKDPRDPEESKEPKEEKQRRR CKPKKPTRRDASPESPSKKGPPIRRH

Table 13 demonstrates the nucleic acid sequence (SEQ ID NO: 191) and amino acid sequence (SEQ ID NO: 192) of GNAS1 from mouse. SEQ ID NO: 192 represents the protein encoded by SEQ ID NO: 191.

TABLE 13

SAGRES TAG#	REF #	SEQ ID#	MOUSE
			SEQUENCE
	191		CCCCCGCGCC CGCCGCGCA TGGGCTGCCT CGGCAACAGT AAGACCGAGG ACCAGCGCAA CGAGGAGAAG GCGCAGCGCG AGGCCAACAA AAAGATCGAG AAGCAGCTGC AGAAGGACAA GCAGGTCTAC CGGGCCACGC ACCGCCTGCT GCTGCTGGGT GCTGGAGAGT CTGGAAAAG CACCATTGTG AAGCAGATGA GGATCCTGCA TGTTAATGGG TTTAACGGAG AGGGCGCGA AGAGGACCCG CAGGCTGCAA GGAGCAACAG CGATGGTGAG AAGGCCACTA AAGTGCAGGA CATCAAAAAC AACCTGAAGG AGGCCATTGA ACCATTGTG GCCGCCATGA GCAACCTGGT GCCCCCTGTG GAGCTGGCCA ACCCTGAGAA CCAGTTCAGA GTGGACTACA TTCTGAGCGT GATGAACGTG CCCGACTTTG ACTTCCCACC TGAATTCTAT GAGCATGCCA AGGCTCTGTG GGAGGATGAG GGAGTGCCTG CCTGCTACGA GCGCTCCAAT GAGTACCAAGC TGATTGACTG TGCCCAAGTAC TTCTGGACA AGATTGATGT GATCAAGCAG GCCGACTACG TGCCAAGTGA CCAGGACCTG CTTCGCTGCC GTGTCCTGAC CTCTGGAATC TTTGAGACCA AGTTCAGGT GGACAAAGTC AACTCCACA TGTCGATGT GGGCGGCCAG CGCGATGAAC GCCGCAAGTG GATCCAGTGC TTCAATGATG TGACTGCCAT CATCTCGTG GTGCCAGCA GCAGCTACAA CATGGTCATT CGGGAGGACA ACCAGACTAA CCGCCTGCAG GAGGCTCTGA ACCTCTCAA GAGCATCTGG ACAACACAGT GGCTGCGCAC CATCTCTGTG ATTCTCTCC TCAACAAGCA AGACCTGCTT GCTGAGAAG TGCTCGCTGG CAAATCGAAG ATTGAGGACT ACTTCCAGA GTTCGCTCGC TACACCACTC CTGAGGATGC GACTCCCGAG CCGGGAGAGG ACCCACCGCT GACCCGGGCC AAGTACTTA TTCGGGATGA GTTCTGAGA ATCAGCACTG CTAGTGGAGA TGGGCGCCAC TACTGCTACC CTCACTTAC CTGCGCCGTG GACACTGAGA ACATCCGCCG TGTCTCAAC GACTGCCGTG ACATCATCCA GCGCATGCAT CTCCCCAAT ACGAGCTGCT CTAAGAAGGG AACACCCAAA TTAAATTCTAG CCTTAAGCAC AATTAATTAA GAGTGAACG TAATTGTACA AGCAGTTGGT CACCCACCAT AGGGCATGAT CAACACCGCA ACCTTCCCTT TTCCCCCAG TGATTCTGAA AAACCCCTCT TCCCTTCAGC TTGCTTAGAT GTTCCAAATT TAGAAGCTT

			MOUSE
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
		192	MGCLGNSKTEDQRNEEKAQREANKKIEKQLQKDKQVYRATHRLI LLGAGESGKSTIVKQMRILHVNGFNGEGGEEDPQAARSNSDGEKATKVQDIKNNLKEA IETIVAAAMSNLVPPVELANPENQFRVDYILSVMVPDFDFPPEFYEHAKALWEDEGVR ACYERSNEYQLIDCAQYFLDKIDVIKQADYVPSDQDLLRCRVLTSGIFETKFQVDKVN FHMFDVGGQRDERRKWIQCFNDVTIAIFVVASSSSYNMIREDNQTNRLQEALNLFKSI WNNRWLRTISVILFLNKQDLLAEKVLAGSKIEDYFPEFARYTTPEDATPEPGEDPRV TRAKYFIRDEFRLISTASGDGRHYCYPHTCAVDTENIRRVFNDCRDIQRMHLPQYE LL

Table 14 demonstrates the nucleic acid sequence (SEQ ID NO: 193) and amino acid sequence (SEQ ID NO: 194) of GNAS1 from human. SEQ ID NO: 194 represents the protein encoded by SEQ ID NO: 193.

TABLE 14

			HUMAN
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
5		193	GCGGGCGTGC TGCCGCCGCT GCGCCGCCG CCGCAGCCCG GCCGCGCCCG GCCGCCGCCG CCGCCGCCAT GGGCTGCCTC GGGAACAGTA AGACCGAGGA CCAGCGCAAC GAGGAGAAGG CGCAGCGTGA GGCCAACAAA AAGATCGAGA AGCAGCTGCA GAAGGACAAG CAGGTCTACC GGGCCACGCA CCGCCTGCTG CTGCTGGGTG CTGGAGAACATC TGGAAAAGC ACCATTGTGA AGCAGATGAG GATCCTGCAT GTTAATGGGT TTAATGGAGA GGCGGGCGAA GAGGACCCGC AGGCTGCAAG GAGCAACAGC GATGGTGAGA AGGCAACCAA AGTGCAGGAC ATCAAAACA ACCTGAAAGA GGCAGATTGAA ACCATTGTGG CCGCCATGAG CAACCTGGTG CCCCCCGTGG AGCTGGCAA CCCCGAGAAC CAGTTCAGAG TGGACTACAT CCTGAGTGTG ATGAACGTGC CTGACTTTGA CTTCCCTCCC GAATTCTATG AGCATGCCAA GGCTCTGTGG GAGGATGAAG GAGTGCCTGC CTGCTACGAA CGCTCCAACG AGTACCAAGCT GATTGACTGT GCCCAAGTACT TCCCTGGACAA GATCGACGTG ATCAAGCAGG CTGACTATGT GCCGAGCGAT CAGGACCTGC TTGCTGCCTG TGCTCTGACT TCTGGAATCT TTGAGACCAA GTTCCAGGTG GACAAAGTC ACTTCCACAT GTTGACGTG GGTGGCCAGC GCGATGAACG CCGCAAGTGG ATCCAGTGTCTCAACGATGT GACTGCCATC ATCTTCGTGG TGGCCAGCAG CAGCTACAAAC ATGGTCATCC GGGAGGACAA CCAGACCAAC CGCCTGCAGG AGGCTCTGAA CCTCTTCAAG AGCATCTGGA ACAACAGATG GCTGCGCACC ATCTCTGTGA TCCTGTTCT CAACAAGCAA GATCTGCTCG CTGAGAAAGT CCTTGTGGG AAATCGAAGA TTGAGGACTA CTTTCCAGAA TTTGCTCGCT ACACTACTCC TGAGGATGCT ACTCCCGAGC CCGGAGAGGA CCCACGCGTG ACCCGGGCCA AGTACTTCAT TCGAGATGAG TTTCTGAGGA TCAGCACTGC CAGTGGAGAT GGGCGTCACT ACTGCTACCC TCATTTCACC TGCGCTGTGG ACACTGAGAA CATCCGCCGT GTGTTCAACG ACTGCCGTGA CATCATTCAAG CGCATGCACC TTCGTCAGTA CGAGCTGCTC TAAGAAGGGAA ACCCCCCAAT TTAATTAAAG CCTTAAGCAC AATTAATTAA AAGTGAACG TAATTGTACA AGCAGTTAAT CACCCACCAT AGGGCATGAT TAACAAAGCA ACCTTTCCCT TCCCCCGAGT GATTTGCGA AACCCCCCTT TCCCTTCAGC TTGCTTAGAT GTTCCAAATT TAGAAAGCTT AAGGCGGGCT ACAGAAAAAG GAAAAAAGGC CACAAAAGTT CCCTCTCACT TTCAGTAAAA ATAATAAAA CAGCAGCAGC AAACAAATAA AATGAAATAA AAGAAACAAA TGAAATAAAT ATTGTGTTGT GCAGCATTAA AAAAATCAA AATAAAAATT AAATGTGAGC

SAGRES TAG#	REF #	SEQ ID#	HUMAN
			SEQUENCE
		194	MGCLGNSKTEDQRNEEKAQREANKKIEKQLQKDKQVYRATHRLL LLGAGESGKSTIVKQMRILHVNGFNGEGGEEDPQAARSNSDGEKATKVQDIKNNLKEA IETIVAAMSNLVPPVELANPENQFRVDYILSVMNVPDFDPPEFYEHAKALWEDEGVR ACYERSNEYQLIDCAQYFLDKIDVIKQADYVPSDQDLLRCRVLTSIFETKFQVDKVN FHMFDVGGQRDERRKWIQCFNDVTAIIFFVASSSYNMVIREDNQTNRQEAQNLFKSI WNNRWLRTISVILFLNKQDLLAEKVLAGSKIEDYFPEFARYTTPEDATPEPGEDPRV TRAKYFIRDEFRLISTASGDGRHYCYPHFTCAVDTENIRRVPNDCRDIIQRMHLRQYE LL

Also suitable for use in the present invention is Genbank Accession No. AJ224868.

A HIPK1 nucleic acid sequence of the invention is depicted in Table 15 as SEQ ID NO. 195. The nucleic acid sequence shown is from mouse.

TABLE 15

5	TAG #	SEQ. ID NO.	SEQUENCE
	S00013	195	CTCCGTNGGGAGCCANCNTGGACGGNGTGTGGGGACCGGTNTCCAGTCNTCTCCGCA AANCGGTCTCCNAGGTGGTTAACCGGNNTTGTTGGTGGNGGTGGGTTTCTTACAGTTA GATGTCANCTCANCTAGTGTGACATCACCCCCAACCGAGTGTGATTTCACCCCCAACAT CCCAATCACATCCCAGCGATTGGGCAGCGCAGGGAGACATTGACTACCTGGGGGATGA CTCTGAGGGTTAGAATTCTCAGTTTACTAAATTGTTGCTGCCATGTCGATTTC AGGGCAGCNAGGGGNATTAGATGCCTCCCTGTCCTTNGA

A contig assembled from the mouse EST database by the National Center for Biotechnology Information (NCBI) having homology with all or parts of a HIPK1 nucleic acid sequence of the invention is depicted in Table 16 as SEQ ID NO. 196. SEQ ID NO. 197 represents the amino acid sequence of a protein encoded by SEQ ID NO. 196.

TABLE 16

SAGRES TAG#	REF #	SEQ ID#	MOUSE
			SEQUENCE
S000013	F3	196	CCGCCACCAAACGCCGGTAAACCACCTCGGAGACTGCTGTGCGGAGAGGACTGGAAACC GGTCCCCACACACTGTCCACGCTGGCTCCCCACGGAGGCCACCCACACCCGCGGGCCGG GCAAGATGCAGTGTACTCAGCCCTCCGCTCTCCGCACCTCCGCTCAGTATGGCTCACA GCTGCAGGTGTTTCGCCCCATCAGTGTGTCGAGTGCCTCTGCAGTGCAAAGAAACTGA AAATAGAGCCCTCTGGCTGGATGTTTCAAGGACAGAGCAGCAACGACAATACTATACCCACA GCAAACCCCTCCAGCTACACAAGGGCAAGCCAGCTCCTCTCACCAAGGTAGCAAATTCAATC TTCCCTGCTTACGACCAGGGCCTCTCTCCAGCTCCTGCCGTGGAGCATATTGTGGTAACAG CTGCTGATAGCTCAGGCAGCGCCGCTACAGCAACCTCAAAGCAGCCAGACCCCTGACTCAC AGGAGCAACGTTCTTGCTTGAGCCATATCAAAATGTGGATTGAAGAGAAAGAGTGAGGAA GTGGAGAGCAACGGTAGCGTGCAGATCATAGAAGAACACCCCCCTCTCATGTCAGAACACAG AACCGTGGTGGGTGCTGCTGCCACGACCACACTGTGACCACCAAGAGTAGCAGTCCAGTG

			MOUSE
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
			GAGAAGGGGATTACCAGCTGGTCCAGCATGAGATCCTTGCTCTATGACCAACAGCTATGAA GTCTGGAGTCCTAGGCCGGGGACATTGGACAGGTGGCAAAGTGTGCTGGAAAGCGGAGCA CCAAGGAAATTGTGGCCATTAAGATCTGAAGAACCAACCCCTCTATGCCAGACAAGGACAGA TTGAAGTGGACATCCTTCCCCTAAGCAGTGAAGAACCAACCCCTCTATGCCAGACAAGGACAGA TTATGAGTGTTCAGCACAAGAACATCACCTGCCTGTGTTGAGATGTTGGAGCAGAACCTT GTACGATTTCTAAAGCAGAACAAAGTTAGCCCCTGCTCAAGTACATAAGACCAATCTG CAGCAGGTGGCCACAGCCCTGATGAAGCTGAAGAGTCTGGTCTGATTGCTGACCTAA ACCTGAAAACATAATGCTAGTCATCCAGTTCGCCAACCCCTACCGAGTGAAGGTATTGACTT TGGTCTGCTAGTCATGTTCAAAGCCGTGTTCAACCTACCTGCAATCACGCTACTACAG AGCTCCTGAAATTATCCTGGATTACCAATTCTGTAAGCTATTGACATGTGGTCACTGGCTGT GTAATAGCTGAGCTGTTCTGGATGGCTCTTATCCTGGTCTCAGAACATACGATCAGATT CGCTATATTCACAAACACAAGGCCCTGCCAGCTGAGTATCTCTCAGTGCCGGAACAAAACA ACCAGGTTTTAACAGAGATCCTAATTGGGGTACCCACTGTGGAGGCTTAAGACACCTG AAGAACATGAATTGAAACTGGAATAAAGTCAAAGAACGCTGGAAAGTACATTTTAAC GTTAGATGACATGGCTCAGTAAATATGTCACAGACTAGAGGGGACAGATATGTTAG CAGAGAAAGCAGATCGGAGAGAGTATATTGATCTCTAAAGAAAATGTCAGGATTGATG CAGATAAGAGAACATCAGCCTCTGAAGACTCTAACCAACCAATTGTGACGATGAGTCACC TCCTGGACTTCTCACAGCAGCCACGTTAAGTCTGTTCCAGAACATGGAGATCTGCA AGCGGAGGGTTACATGTATGACACAGTCAGTCAGATCAAGAGTCCCTCACTACACATG TCGCTCAAATACAAGCACAAATCTAACCATGAGCTCAGCAACCAGCTAACACAGTGC ACAATCAGGCCAGTGTCTAGCTTCCAGCTCTACTGCAGCAGCAGCTACCCCTCTGG CTAATTCAAGATGTCTCGCTGCTAAACTACCAATCGGTTGTACCCATCGTCGGCAGCGC CAGTCTGGAGTTGCCAGCAGGGTGTCTTACAACCTGGAACCAACCCAGATCTGCA CTCAGACAGATCCATTCCAGCAAACATTATAGTATGCCACCTGCTTCAACTGGAC TACAAGCAACAAACAAAGCATTCTGGATTCCCTGTGAGGATGGATAATGCTGTGCAATTG TACCCAGGCCCTGCTGCTCAGCCGCTGAGATCCAGTCAGGAGTACTCACACAGGGAA GCTGTACACCAACTATGGTAGCAACTCTCCACCCCTCAAGTAGCCACCATCAGGCCAGT ATGCGGTGCCCTTACCCCTGAGCTGCCAGCAGGCCGGGGCGCTGGTGAACAGACTG CTGCTGTACTGCAAGCCTGGCCTGGAGGAACCCAAACAAATTCTCTGCCCTCAGCCTGG AGCAGCTGCCGGGGTAGCTCTGCACAACCTGTCCAGCCTGCTGAGTGAATTCCAGAGG CCATGGGAGCAGCAAACAGCTAGCTGACTGGAGGAATGCCACTCTCATGGCAACCA ACAGCACTATTATGCAGCAGCCATCTTGCTGACCAACCATGTCACCTGGCCACTGCTC AGCCTCTGAATGTTGGTGTGCCCCTGAGACAAACACAGTCTAGTCCCTCCCTT CAAAGAAGAATAAGCAGTCTGCTCAGTTCTACCAATCCTCTGGAAGTCTGCTGCC CTCAAGTTATTCTCTGGTGGAGTAGTCCTCTCGTACCCATCTTCTATAATTCCC TAGTCTCTGCTCAAGACCAGCATGCCAATCATCATTCCAGATACCCCGGCCCTCTG TGAGTGTCTACTATCCGTAGTGACACTGATGAAGAAGAGGAACAAATACAAGCCA ATAGCTCGAGCCTGAAGGCAGGTCTAATGTCATCAGTTATGTCACTGTCATGATTCTC CAGACTCTGACTCCTCCCTGAGCAGCCCACATCCCACAGACACTCTGAGTGTCTGCC GCAACAGTGGGACCCCTGGAGGGACCTGGCAGACCTGCTGAGCAGATGGCATTGGCACCC GTACTATCATTGTGCCCTCTTGAACACAGCTGGCAGACTGCACTGAGCAACACAGG CCTCAGGTCTCTTAGCAGTAAGACCAAGCCAGTGGCCTCAGTGAGTGGCAGTCATCTG GATGCTGTATCACTCCACGGGGTACCGGGCTCAGCGAGGGGGAGCCAGCGCGGTGCA CACTCAACCTTAGCCAGAACAGCAGTCAGCTGCACTGCTCAACCTCGCAGGAAAGAAGCA GCAACCCCTGCTCCCCGAGACAGCAGGCAATTGTGGCCCCGCTCTCCAAGCCCCCTACG CCTCCAGCATGGCAGCCACTGCACTGACGGGGCACCAACTTGGCCCCAGCCCCCTG

SAGRES TAG#	REF #	SEQ ID#	MOUSE
			SEQUENCE
			CTCACCTGCCAAGCCAGCCTCACCTGTATACGTACGCTGCCCCACTTCTGCTGCTGCAT TGGGCTCCACCAGTCCATTGCTCATCTGTTCTCCCCCAGGGTCTCAAGGCATGCTG CAGCTTATACCACACACCCCTAGCACTCTGGTGCACTCAGGTTCTGTCAAGTGTGCGGGCCA GCCTCCTCACTTCTGCCAGTGTGGCCCTGCTCAGTACCAACACCAGTTGCCACTCAGT CCTACATCGGGTCTTCCGAGGCTAACAAATTACACTGGATACCCGCTGAGTCTTACCA AGATCAGTCAGTATTCTACTTGTAGTTGATGAGCACGAGGAGGGCTCCGTGGCTGCCTG CTAAGTAGCCCTGAGTTCTTAATGGGCTCTGGAGAGCACCTCCATTATCTCCTCTTGA GTTCTAGCCAGCAGCGTTCTGCCGGGCCACTGAAGCAGAAGGCTTTCCCTGGGAA CAGCTCTCGGTGTTGACTGCATTGTTGCACTCCTCCAAAGTCTGCCCTGTTTTTAATT TTTATTCTGTGACAGCATTTGGACGTTGGAAGAGCTCAGAAGCCCCTTGCAGT TACCAAGGAAGAAAGATCGTCTGAAGTTACCCCTGTCTACATTTGGTCTCTTGA TGGTTCTATAAAATGTTTTAAAATGAAGTAAAGCTCTTCTTACGAGGGAAATGCTGA CTTGAATCCTGAGCAGATGAGAAAGAGTCATTACTTTTGTGCTTAAAGAAACTAA ACACAAGACTCCTGCTTTATTTGAAAGCAGCTTAGCAAGGGTGTGCTTATGGCGT ATGGAACACGAATGATTCTATTTGATGTCGTGCTGCCTTACTGGCAGTTGTTAGAGT TTTAGTACAACGAGTCAGTGAACCTGTGCACTGCTGCTGAGCTGCTCGCAGAGCAGCA CTGAACAGGCAGCCAGCGCTGCTGGAGGAAGGTGAGGGTGGAGGACTGTGCCACCAGG ATTCAATTCTAAATGAAGACCATGAGTTCAAGTCTCCCTCTCTAGTTAACTTAA TTCTCCTTATAGAAAAGCCAGTGAGGTGGTAAGTGTATGGTGGTGGTTGCATAAATAG TATGCAAATCTCTCTAGAATGAGACTGAGCACTGATAAAACATTGCCATAAGATTCT ATGAATTCTAAATAACACGTCTGTGTTTCTCATCTCTCCCTCTGTTCTATGTGACT TATTGAGGGAAAACAAAGAAACTAAAACCAAGATAAGTTGTATAGTTTATACTT TAAAGTAGCTTCTTGTATGCCAACAGCAAATTGAATGCTCTTACTAAGACTTATGT AATAAGTCATGTAGGAATTGCAAGAAAATTTAAAGTTTATTACTGAATTAAAAT ATTTAGAAGTTTGTATGGTGGTTAAATATTGCTATAATTAAATATGTACATAT TGATTAGAAGAAAATAACAATTCTCTAACCCAAAATGTTATTGTAATCAAATGT GTAGTGATTACACTGAATTGTGTTAGTGCTGAGCTGCTGAGAGAGCAGGGCATT AGATGGATTATGCTCCATTGTATTAAACCAAAATGAACGTGATACTTGTGGATGT GTGAACTAATTGCAATTCTATTAGAGCATATTACTGTAGTGCTGAGAGAGCAGGGCATT GCCTGCAGAGAGGAGACCTTGGGATTGTTGACAGGTGTGCTGGTGGAGGAGTTGTC AGTGTGTCTTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCT ATGTAGTGGTTAATAGAGTTACAGTGAGCTGCCCTAGGATGACCAGCAAGCCCCAGTG ACCCCAAGCTGTCGCTGGGATTTAACAGAGCAGGTTGAGTAGCTGTTGTGAAATGC GTTCTGTTCTCAGTCTCCCTACCGACAGTGACAAGTCAAAGCCGAGCTTCTCTCT ACTGCCACCTCTGCCCCGTTCCATTGGATCTCAGCTCAGTCTCACAGAAGCATTCC CTAACGTGGCTCTCTCACTGTGCCCTGCTACCTGGCTCTGTGAGAGGTTCAAGGAAGCAGG CGAGAAGAGTGACGCCAGTGCTAAATATGCATATTGAAGGTTGTGCTTACTTAGGGT GGGATTCTTCTCTCTCCATGTGATATGATAGTCCTCTGCTAGCTGTCGTTCC TGGTAAACTTGTGTTGGTTTTTTTTTTTTGTTGTTGTTTTTTAAAGCATGTA CAGATGTGTTATACCAAGAGCCTGTTGATTGCTTAATATGTCCCCTACTACGAGAAG GGTTTGAGAAGTACTGGTGACAAGAAGCTCACAGAAAGGTTCTTAATTAGTGACGAA TATGAAAAAGAAAGCAAACCTTGTGATCTGAACAATTCTGAGGTTCTTGGGACAA CATGTTGTTCTGGGGCCCTGCACACTGTAAAATTGCTCTAGTATTCAACCCCTCCATGG ATTTGGGTCAAGTTGAAGGTACTAGGGGTGGGACATTCTGCCCATGAGGGATTG GGAGAAGGTTAACCTAAGCTACAGAGTGGTCCACCTGAATTAAATTATCAGAGTGG AATTCTAGGATTGGTCTGTTAGGTGGTCAAGGAGGTGCAAGGATGGAGATGGGAGATT

			MOUSE
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
			TCATGGAACCGTTCAGGAAAGCTCTGAACCAGGTGGAACACCGAGGGCTGTCACCGAA CTTGGAGTTCTCATCATGGGAGGAAGAGTTCCAGGGCAGGGCAGGTAGTCAGTTA GCCTGCCGGCAACGTGGTGTGTTCTTCTTAATCATTATAAGCTGTGCGTT CAGCAGTCTGTTGGTGGAGATAACCACGCATCATTGTGTAGTTGACTAGTGTATAC CGTTATGTCAATTGTGTGATCTTGTGTTCTTCCCCAAGCATTCTGGGTTTT TCCTATTAAATACAGTTCTAGGCAAACATTAAAAACCTTTCTCTATAA GGGACAAGATTATTGTTTATAGGAATGAGATGCAGGGAAAAAACAAACCAACCCGT CCCCACTCCTCACCTCCCTAATCCAATAAGCAGTTATTGAAGATGGGAGTCTAAATTAA TGGGAAAAGAGGATGCCTAGGAGTTGCATCGTTACCTGAGACATCTGGCTAGCAGTGT ACTTTACAGACTTGAGGTTGTCACTCTGCAAACGTGACATTGAGATTCTAGATAAC CCATCTGTGTCGCTGAATGTGTGCGCCAGACATAGTTACATTCTGGCCTGG GGCTTAACATTGACTGCTGCCCTGATGGCATGGAGAGCCCTACGAACATAGCGCTG ACTAGGTCACTGCCTGACCTTGAACAGCTTAAGGCTTAAACCTTCTCTAGAACG TGCATTCCAGTTCTCCCTCCAGGTGAGAGAGGAACGGAACTGGAAGGGTTGCATAGGACA CACCAGGACACTTAGTCACTCCAGAGTCCCCAGTTGCAACTAGGAGGTGGTACCTGTT AACCCCAGGAAGAACCCCATTCAACAGTTCCGGCCATTGAGAGCCTGCTTTGTT GTTGCTCATCCGTACATCCGCTAGAGGGGCTTAGCCAGGCCAGCACAGTACTGGCTGT CCTATTCTGCATTAGTATGCAGGAATTACTAGTTGAGATGGTTGTTAGGATAGGAG ATGAAATTGCCCTCGGTGACAGGAATGCCAAGCCTGCTTGTGTTTTTTAAATGA TGGATGGTGCAGCATGTTCCAAGTTCCATGGTTGTTGCTAAATTATATAATG TGTGGTTCAATTCAATTCAATTGCTTGGAAAATAATTCACTATATGTAGCAGTACATTATA TGTACATTATATGTAATGTTAGTATTGCTTGAATCCTGATATTGCAATGGAATT CTAATTATTAAATGTATTGATATGCTAAAAAA
	197		MASQLQVFSPPSVSSAFCSAKKLKIEPSGWDVSGQSSNDKYYTHSKTLPATQQQASSSHQVAN FNLPAYDQGLLLPAPAVEHIVVTAADSSGSAATATFQSSQTLHRSNVSLEPYQKCGLKRKSEEV ESNGSVQIIEHPPLMLQNRTV/GAAATTTVTCKSSSSSGEHDYQLVQHEILCSMTNSYEVLEFL GRGTGQVAKCWKRSTKEIVAIKILKNHPSYARQGQIEVSILSRLSSENADEYNFVRSYECFHKN HTCLVFEMLEQONLYDFLKQNKFSPPLPKYIRPILQQVATALMKLKSLGLIADLKPENIMLVDPVRQ PYRVKVIDFGSASHVSKAVCSTYLQSRYYRAPEIIGLPFCEAIDMWSLGCVIAELFLGWPLYPGAS EYDQIRYISQTQGLPAEYLLSAGTKTRFFNRPNLGYPLWRLKTPPEEHELETGIKSKEARKYIFNC LDDMAQVNMSSTDLEGTDMLAEKADRREYIDLKKMLTIDADKRITPLKTLNHQFVTMSHLLDFPHS SHVKSCFQNMEICKRRVHMYDTVSQIKSPFTTHVAPNTSTNLMSFSNQLNTVHNQASVLASSST AAAATLSLANSDVSLNYQSALYPSSAAPVPGVAQQGVSLQPQTTQICQTDPFQQTFIVCPFAQ TGLQATTKHSGFPVRMDNAVPIVPQAPAAQPLQIQSGVLTQGSCTPLMVATLHPQVATTPQYAV PFTLSCAAGRPAVEQTAAVLQAWPGTQQILPSAWQQLPGVALHNSVQPAAVIPEAMGSSQQ LADWRNAHSHGNQYSTIMQQPSLLTNHVTLATAQPLNVGVVAHVRQQSSSLPSKKNQKQAPVS SKSSLEVLPQSQVYSLVGSSPLRTTSSYNSLVPVQDQHQPIIPDTPSPPVSVITIRSDTDEEEEDNKYK PNSSLKARSNVISYTVNDSPDSDSLSSPHPTDTLSALRGNSGTLEGPGRPAADGIGTRTIIVP PLKTQLGDCTVATQASGLSSKTPVASVSGQSSGCCITPTGYRAQRGGASAVQPLNLSQNQQS SSAISTSQERSSNPAPRQQAFVPLSQAPYAFQHGSPHLHSTGHPHLAPAPAHLPSPQPHLYTYAA PTSAAALGSTSSIAHLFSPQGSSRHAAAYTTHPSTLVHQPVSVGPSLLTSASVAPAQYQHQFAT QSYIGSSRGSTIYTGYPLSPTKISQYSYL

15 Also suitable for use in the present invention is the sequence provided in Genbank Accession No. AF077658.

A contig assembled from the human EST database by the NCBI having homology with all or parts of a HIPK1 nucleic acid sequence of the invention is depicted in Table 17 as SEQ ID NO. 198. SEQ ID NO. 199 depicts the amino acid sequence of an open reading frame of SEQ ID NO. 198 which encodes the C-terminal portion of human HIPK1 protein.

5

TABLE 17

SAGRES TAG#	REF #	SEQ ID#	HUMAN
			SEQUENCE
S000013	F30	198	CACACCGCAGTATGCCGCTTACTCTGAGCTGCCAGCCGGCCGGCGCTGGT TGAACAGACTGCCGCTGACTGGCGTGGCCTGGAGGGACTCAGCAAATTCTCCTGCCTC AACTTGGCAACAGTTGCCCTGGGGTAGCTCTACACAACCTGTCCAGCCCACAGCAATGAT TCCAGAGGCCATGGGGAGTGGACAGCAGCTAGCTGACTGGAGGAATGCCACTCTCATGG CAACCACTACAGCACTATCATGCAGCAGCCATCCTGCTGACTAACCATGTGACATTGGC CACTGCTCAGCCTCTGAATGTTGGTGTGCCATGTTGTCAGACAACAATCCAGTC CCTCCCTCGAAGAAGATAAGCAGTCAGCTCCAGTCTCTCCAAGTCCCTCTAGATGT TCTGCCCTCCCAAGTCTATTCTCTGGTTGGGAGCAGTCCCTCCGCACCACATCTCTTA TAATTCCCTGGTCCCTGTCCAAGATCAGCATCAGCCCATCATCATTCCAGATACTCCAG CCCTCCTGTGAGTGTCACTATCCGAAGTGACACTGATGAGGAAGAGGACAACAAATA CAAGCCCAGTAGCTCTGGACTGAAGCCAAGGTCTAATGTCATCAGTTATGTCACTGTCAA TGATTCTCCAGACTCTGACTCTTCTTGAGCAGCCCTTATTCCACTGATAACCTGAGTGC TCTCCGAGGCAATAGTGGATCCGTTTGGAGGGGCTGGCAGAGTTGTCAGATGGCAC TGGCACCCGCACTATCATTGCCCCACTGAAAACCTCAGCTTGGTGAUTGCACGTGAG AACCCAGGCCTCAGGTCTCCTGAGCAATAAGACTAAGCCAGTCGCTCAGTGAGTGGCA GTCATCTGGATGCTGTATCACCCCCACAGGGTATCGAGCTCAACGCCGGGGGACAGTGC AGCACAACCAACTCAATCTTAGCCAGAACCGAGCAGTCATGGCGGCTCAACCTCACAGGA GAGAAGCAGCAACCCAGCCCCCGCAGCAGCAGCGTTGTGGCCCTCTCTCCAAAGC CCCCCTACACCTCCAGCATGGCAGCCGCTACACTGCACAGGGCACCCACACCTGCCCC GGCCCTGCTCACCTGCCAAGCCAGGCTCATCTGTATACGTATGTCCTCCGACTCTGC TGCTGCACTGGGCTCAACCAGCTCCATTGCTCATCTTCTCCCCACAGGGTCTCAAG GCATGCTGCAGCCTATACCAACTCACCTAGCATTGGTGCACCGAGTCCCTGTCAGTGT TGGGCCAGCCTCTCACCTGCCAGCGTGGCCCTGCTCAGTACCAACACCAGTTGC CACCAATCCTACATTGGCTTCCCGAGGCTAACAAATTACACTGGATAACCGCTGAG TCCTACCAAGATCAGCCAGTATTCTACTTATAGTTGGTGAUTGAGCATGAGGGAGGAGGAATC ATGGCTACCTCTCCTGGCCCTGCGTTCTTAATATTGGCTATGGAGAGATCCTCTTITA CCCTCTGAAATTCTAGCCAGCACTGTTCTGCAGGGGCCACTGAAGCAGAAGGTT TTCTCTGGGGAACCTGCTCAGTGTGACTGCATTGTTGAGTCTCCCAAAGTTGC CCTATTTAAATTCTTGTGACAGTAATTGGTACTTGGAAAGAGTTAGATG CCCACCTCTGCAGTTACCAAGGAAGAGAGATTGTTCTGAAGTACCTCTGAAAAATAT TTTGCTCTGACTTGATTCTATAATGCTTTAAAAACAAGTGAAGCCCTCTTTAT TTCATTTGTGTTATTGTGATTGCTGGTCAGGAAAAATGCTGATAGAAGGAGTTGAAATC TGATGACAAAAAAAGAAAAATTACTTTTGTGTTATAAAACTCAGACTTGCCTATTT ATTTAAAAGCGGCTTACACAATCTCCCTTTGTTATTGGACATTAAACTACAGAGT TTCAGTTTGTGTTAATGTCATATTACTTAATGGCAATTGTTATTGCAAACACTG GTTACGTATTACTCTGTGTTACTATTGAGATTCTCAATTGCTCCGTGTTGTTATA AGTAGTGTGTTAAAAGGCAGCTCACCATGGCTGTACTTAATGTGAGGAGAATCCATATC TGCCTGAAAACACCAAGTATTCTTTAAATGAAGCACCAGTGAATTCTTTAAATTAT

HUMAN			
SAGRES TAG#	REF #	SEQ ID#	SEQUENCE
			TTTTAAAAGTCTTCTCTCTGATTCACTAAATTTTTATCGAAAAAGCCATTAA GGTGGTTATTATTACATGGTGGTGGGGTTTATTATATGAAAATCTGTCTATTATG AGATACTGGCATTGATGAGCTTGCTAAAGATTAGTATGAATTTCAGTAATACACCTC TGTTTGCTCATCTCCCTCTGTTTATGTGATTGTTGGGGAGAAAGCTAAAAAA CCTGAAACCAGATAAGAACATTCTTGTGTATAGCTTTACTTCAGAAGTAGCTTCCCT TGTATGCCAGCAGCAAATTGAATGCTCTTATTAAAGACTTATATAATAAGTCATGTAG GAATTGCAAAAAAATTTTAAATTACTGAATTAAATATTAGAAGTTTG TAATGGTGGTGTAAATATTACATAATTAAATATGTACATATTGATTAGAAAAAT AACAAAGCAATTTCCTGCTAACCCAAATGTTATTGTAATCAAATGTGTAGTGTAC ACTTGAATTGTGTACTTAGTGTATGTGATCCTCCAGTGTATCCGGAGATGGATTGA TGTCTCCATTGTATTAAACCAAAATGAACTGATACTTGTGGAATGTATGTGAACTAAT TGCAATTATATTAGACATATTACTGTAGTGTGAATGAGCAGGGCATTGCCTGCAAGG AGAGGAGACCCCTTGGAAATTGTTGCACAGGTGTCTGGTAGGGAGTTTCAGTGTG GTCTCTCCCTCCCTTCTCCTCCCTTATTGTAGTGCCTTATATGATAATGTAGT GGTTAATAGAGTTACAGTGAGCTGCCTAGGAATGGACAGCAAGCCCCCGTGGACCC AAGTTGTTACCCGGATTATCAGAACAGGATTAGTAGCTGTATTGTGAAATGCATTGTT CTCAGTTCCCTGCCAACATTGAAAATAAAACAGCAGCTTCTCCCTTACCAAC TCTACCCCTTCCATTGGATTCTCGGCTGAGTTCTCACAGAACGATTCCCCATGTG GCTCTCTCACTGTGCGTTGCTACCTGCTCTGTGAGAACATTAGGAAGCAGGTGAGAGGA GTCAAGCCAATATTAAATATGCAATTAAAGTATGTGCAATCACTTTAGAATGAAT TTTTTTCTTCCATGTGGCAGTCCTCCTGCACATAGTTGACATTCTAGTAAAA TATTGCTTGTGAAAAAAACATGTTAACAGATGTGTTATACAAAGAGCCTGTTGAT TGCTTACCATGCCCCATACTATGAGGAGAAGTTGTGGCCTGGTACAAGGAAC TCACAGAAAGGTTCTAGCTGGTGAAGAATATAGAGAAGGAACCAAGCCTGTTGAGTC ATTGAGGCTTTGAGGTTCTTAAACAGCTTGTATAGTCTGGGGCCCTCAAGCTG TGAAAATTGCTTGTACTCTCAGCTCTGCATGGATCTGGTCAAGTAGAAGGTACTGG GATGGGGACATTCTGCCATAAAGGATTGGGGAAAGAAGATAATCCTAAATACAGG TGTGTTCCATCGAATTGAAATGATATTGAGATATAATTAGGACTGGTTCTGTG TAGATAGAGATGGTGTCAAGGAGGTGCAGGATGGAGATGGGAGATTTCATGGAGCCTGG CAGCCAGCTCTGTACCAAGGTTGAACACCGAGGAGCTGTCAAAGTATTGGAGTTCTCA TTGTAAGGAGTAAGGCTTCCAAGATGGGCAGGTAGTCGTACAGCCTACAGGAACAT GTTGTGTTCTTATTAAATCATTATATTGAGTTGTTTACGCACTATATTG GTCAAGATAGCCAAGCAGTTGTATAATTCTGCACTAGTGTCAACAGTTCTGGTC AACATGTGTGATCTTGTGCTCTCTTGTGAAAGCACATTCTGATTCTGTTGGAAC ACAGGTCTAGTTCTAAAGGACAAATTGTTGCTCTGTCTTTCTGTAAGGGACAA GATTGTTGTTTGTAAAGAAATGAGATGCAGGAAAGAAAACCAATCCATTCTGCAC CCCAGTCCAATAAGCAGACCAACTTAAGATAGGAGTCTAAACTCCACAGAAAAGGATAA TACCAAGAGCTGTATTGTTACCTAGTCAGTGCCTAGCAGTGTGGCTTAAACT AGAGATTTCAGTCTAGTGTGCAAACCTGGCATTCCGATTTCAGCATAAAATCCA CCTGTGCTGCTGAATGTGATGTGCTACTGTGGCTTAGATTCTGCTGGGG TAGCCCTGTTGGCCCTGACAGGAAGGGAGGAAGCCTGGTGAATTAGTGAAGCAGCTGG CTGGGTACAGTGACCTGACCTCAAACCGAGCTTAAGGCTTAAAGTCTCTCAGAACTT GGCATTCTCAAACCTCTCCCTTCCGGGTGAGAGAAGAAGCAGGAGAAGGGTTAGTGTAGC CACTCTGGCTCATAGGGACACTGGTCACTCAGAGTTTAATAGCTCCCAGGAGGTG ATATTATTTCACTGCTCAGCTGAAATACCAACCCAGGAATAAGAACTCCATTCAAAC AGTTCTGGCCATTCTGAGCCTGCTTTGTGATTGCTCATCCATTGTCTCCACTAGAGGG

SAGRES TAG#	REF #	SEQ ID#	HUMAN
			SEQUENCE
			GCTAAGCTTGAUTGCCCTTAGCCAGGCAGCACAGTAATGTGTGTTGTTCA TATGCAAAAAATTCACTAGTTGAGATGGTTGTTAGGATAGGAAATGAAATTGCTC AGTGACAGGAGTGGCCCGAGCCTGCTTCCTATTTGATTTTTTTTTAACTGATAG ATGGTGCAGCATGTCTACATGGTTGTTGCTAAACTTATATAATGTGTGGTTCAA TTCAGCTTGGAAAATAATCTCACTACATGTAGCAGTACATTATATGACATTATGTAA TGTTAGTATTCTGCTTGAATCCTGATATTGCAATGGAATTCTACTTTATTAAATGT ATTGATATGCTAGTTATTGTCGATTAAACTTTTGCTTCTCCCTTTGG TTGTGCGCTTCTTTACAACAAGCCTCTAGAAACAGATAGTTCTGAGAATTACTGAGC TATGTTGTAATGCAGATGTACTTAGGGAGTATGTAAAATAATCATTAAACAAAGAAA TAGATATTAAATTAATACTAACTATGGAAAAGGGTCCATTGTGAAAACATAGTT ATCTTGGATTCAATGTTGCTTGGTTTACAAAGTAGCTTGTATTTCAGTATTTC TACATAATGGTAAATGTAGAGCAATTGCAATGCATCAATAAATGGTAAATTTC
	199		TPQYAVPFTLSCAAGRPALEQTAAVLAWPGBTQQILLPSTWQQLPGVALHNSVQPTAMPEAMG SGQLADWRNAHSHGNQYSTIMQQPSLLTNHVTLATAQPLNVGVAHVVRQQQSSLPSKKNQ APVSSKSSLDVPSQVSLVGSSPLRTTSSYNSLVPVQDQHQPIIPDTPSPPVSVITRSDTEEEE NKYPSSSGLKPRSNVVISYVTVDSPSDSSLSSPYSTDLSALRGNSGSVLEGPGRVVADGTGR TIIVPLKTQLGDCTVATQASGLLSNKTGPVASVSGQSSGCCPTGYRAQRGGTSAAQPLNLSQN QQSSAAPTSQERSSNPAPRRQQAFVAPLSQAPYTFQHGSPHLHSTGHPHLAPAPAHLPSQAHLTY AAPTSAAALGSTSSIAHLFSPQGSSRHAAYTTHPSTLVHQVPVSVGPSLLTSASVAPAQYQHQFA TQSYYGSSRGSTIYTGYPLSPTKISQSYL

The JAK1 nucleic acid sequences of the invention are depicted in Tables 18 and 19. The nucleic acid sequence shown in Table 18 is from mouse. The nucleic acid sequence shown in Table 19 is from human. The nucleic acid sequence shown in Table 22 is Sagres Tag No. S00039. The JAK1 amino acid sequences are shown in Tables 20 and 21. Table 20 shows the amino acid sequence from mouse and Table 21 shows the amino acid sequence from human.

Table 18 : JAK1 Nucleotide Sequence from Mouse

Sagres Tag No. S00039	Seq. ID No. 200	CAGCCGGAGTAGCCGGCAGCCGCTGACGCCCGCGGGTCCGCCCAAGCCTCGTCGCTT TCGGTGCCCTCCCTAGCCGGGTGTCACGCCGGACCCCTGCACGGCAGGCTGAGTTGCCTG CAGACTCTGACCCAGATCGACCCCTGCAGCCAAGGAGCCGGCGGGCGCACACGGAACTG ATCAGCTCTGAATGGGCTTGAAGGTAAGAAGAAAAATCCAGTCTGCTTCAGGGACACTGGAC AACCGAATAATGCAGTATCTAAATATAAAAGAGGACTGCAATGCCATGGCCTCTGTGCTAAAAT GAGGAGCTTCAAGAAGACTGAGGTGAAGCAGGTGGCTGAGCTGGAGTGGAGGTGACTTTC TATCTGTTGGACAGGGAGCCCTCCGCTGGCAGCGGAGAGTATACAGCCGAGGAGCTGTGCA TCAGGGCCGCCAGGGAGTGCAGTATCTCTCTCTGTCACAACACTCTTCGCCCTGTACGATGAG AGCACAAGCTCTGGTACGCTCCGAACCGAATCATCACTGTGGATGACAAAACGTCTCCGGCT CCACTACCGCATAGGGTCTACTTACCAACTGGCACGGAAACCAATGACAACGAACAGTCTGTATG GCGACATTCTCAAAGAAGCAGAAAAACGGCTATGAGAAGAAAAGGGTCCAGAAGCAACCCAC TCCTTGATGCCAGTTCACTGGAGTATCTGTTGCACAGGGACAGTATGATTGATCAAATGCCCTGG CTCCCATTGGGACCCCAAGACGGAGCAAGACGGACATGATATTGAAAATGAGTGCTGGCATG GCGGTCTGGCCATCTCCCACTATGCCATGTGAAGAAGAGTGCAGTTGCCGAACCTCCAAAAGA CATCAAGCTACAAGCGATATACTCCAGAAACATTCACTGAAATAATCCATCAGACAGAGGAACCTTCA AGGATGCGAATAAAATAATGTTTCAAGGATTCTGAAGGAATTAAACAACAAGACCATCTGTGACA GCAGTGTGATGACCTGAAGGTGAAAATACCTGGCTACCTTGGAAACTTCTACATTGACAAAACATT ATGGAGCTGAAATATTGAGACTTCTATGCTACTGATTTCATCAGAAATGAAATTGAGTCGATGCCA TTCGAATGACAGTGGCAATGTTCTATGAGGTACTGGTACTGAGGAAATCTGGGATCCAGTGGC GGCAGAAACCAAATGTTCTCTGTGAAAGGAAAATAAAACTGAAGCGGAAAAAAACTGGAAAT ATAATAACAAAGAAGGATGATGAGAGAAACAAACTCCGGGAAGAGTGGAAACAATTCTTCTT CCCTGAAATCCCCACATTGTAATAAAGGAGTCTGTGGTCAGCATTAAACAAACAGGACAACAAAAA CATGGAACCTCAAGCTCTTCTCGAGAGGAAGCCTTGTCTTGTGCTCCCTGGATGGCTACTT CCGGCTCACTGCAAGATGCCACCATTACCTCTGTACTGATGGCTCCCTGGATGGCTACT TATACAGAACGGCTGCCACGGTCAAATGTCACAGAATATGCCATCAATAAGCTGGCAGGAAG GGAGTGAAGAGGGGGATGTCGTGAGGTGGAGGCTGCACCGACTTGAACAACATTCTATGACT GTCACCTGCTTGTGAAAGTCTGAGGTATTGGTGGCCAGAACGAGTTCAAGAACTTCA GTACAGAAGGGCCGCTACAGCCTGCATGGCTCATGGACCACCTTCCAGCCTGCCAGACCTCAT GAACCAACCTCAAGAAGCAGATCCTGCCACGGACACATAAGCTTGTGCTGAAACGATGCTG AGCCTAAAGCCTCGAGAAATCTCACTGTGTAAGCCACTAAGAAAAGCCAGGGAGTGGCAGCCT GTCTACTCCATGACCCAGTGCAGGTTGAGGTTGATCGGATCCTTAAGAAAAGATATTATAACAAGGTGAGC CTTGGCAGAGGCAACAAACATATCTATTCTGGGACCCCTGCTGGACTACAAGGATGAGGAAG AATTGCTGAAGAGAAGAAGATAAAAGTGTACCTCAAAGTCTAGACCCCAGCCACGGGACATCTC TCTGGCTTCTTGAGGCTGCTAGCATGATGAGACAGGTTCCACAAACATATAGTGTACCTCTA CGCGTGTGTGTCGAGATGTTGAAAGGTTGAGGGGGGGGGCGTTG GATCTCTCATGACCCGGAAAGTGTGCGCTTACTACCCCTGGAAAGTCAAGGTTGCCAACAG CTGGCCAGTCCCTGAGTTACTTGGAAAGATAAACCTGGTGTGATGAAAGTGTGACTAAAAAC CTCTTCTGGCCCGTGGGGCATTGACAGTGACATTGGCCCGTTCATCAAGCTTAGTGTGACCTGG CATCCCAGTCTGTGCTGACCAGGAAGAGTGCATAGAGCGAATCCCTGGATCGCTCTGAGT GTGTTGAAGACTCCAAGAACCTGAGTGTTGAGGTTGAGCTTGGAGCTTGGAAACACGCTCTGG GAAATCTGCTACAACGGAGAGATTCTCTCAAAGAACAGACCCCTGATTGAGAAGAGGGTTTAT GAAAGCCGTCGAGGCGATTGACAGTGACATTGGCCCGTTCATCAAGCTTAGTGTGACCTGG GAACATGACCCCAACAGAGACCCCTTCTCCGAGGACATCATGAGGGACATTAACAAGCTGGAGG AGCAGAACATTGAGACATTGTTAGAAAAGCAGCCAACAACAGAGGTGGACCCACTCACTT AGCGGTTCTGAAGAGGATTGTCAGTTGGAGAGGGTCACCTTGGGAGGGTGAAGACTCTGAGA TATGATCTGAGGGAGAACACAGGGGAGCAGGTAGTGTCAAGTCCCTGAGGCTGAGAGTGT GAGGTAACACATAGCTGATCTGAAGAAGGAGATAGAGATCTACGGAAACCTTACCATGAGAAC TTGTGAAGTACAAGGAATCTGATGAGAACAGGGCAATGGTATCAAGCTCATGAGGTT TGCCTTCTGGGAAAGCTTAAAGGAGTATCTGCCAAAGAATAAGAACAAAATCAACCTCAAACAGC TAAAATGCCATCCAGATTGTAAGGGAGGGACTACTTGGGTTCTGCCAATACGTTACCGGG ACTTAGCAGCAAGAAATGTCCTGTTGAGAGTGAGCATCAAGTGAAGATCGGAGACTTGGTTAA CCAAAGCAATTGAAACCGATAAGGAGTACTACACAGTCAGGACGACCCGGGACAGCCAGTGG TGGTACGCTCCGGAAATCTTAAAGGAGTATCTGCCCTGAGGTTTATCAGCTTATGAGAAAATG TGACACTGACAGGCTGCTACTTGACTCAGATTTAGTCCCATGGCCCTGGCTCTGAGGTT TGATAGGCCCACCTCATGGCCAGATGACAGTGACACGGCTTGTGAAGACTCTGAAAGAAGGAAAG CGTCTGCCATGTCACCCAACTGTCCTGATGAGGTTTATCAGCTTATGAGAAAATG CAACCATCTAACCGGACAACCTTCAAGACCTTATTGAGGGATTGAGGACTTAAAGAAG CATGAACAACATTAAATTCCCTTAAATCAACATCTCCCTGCCAAGCCATTAAAACGTTTAA GTGAAAAGTTGATTCTGCCCTAAAGTCTCAACAAACATCGAGTTACACATATGCTATGTC ACACTGCACTCAGTGTGAGATGCTATGTCACACTGTCACTGAGTGTGGAACTTCTCTT AAAGGTGTAACATCTTAAATTGGTGTGATGAAATGAGACACCAAAAGACTAGATTG TCCTCTGGAAACAACCGAATGTCAGCTGATGAGGACTGTGCCCTGGCATATTGATCTCA GATAAAAACCTTGGAACCTGGACTTGCTGACACTCTCCCTGCCCTGAAATCTCA GTACAAGCACGTAAGACCACTTAGTACTATTGTTCTATTAAAAAA
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Table 19: JAK1 Nucleotide Sequence from Human

Sagres	Seq. ID	
Tag	No.	
No.		
S00039	201	TCCAGTTGCTTGGAGAACACTGGACAGCTGAATAATGCAGTATCTAAATATAAAAGAGGACTGC AATGCCATGGCTTCTGTCTAAATGAGGAGCTCCAAGAAGACTGAGGTGAACCTGGAGGCCCTGA GCCAGGGTGGAAAGTGTCTTCTATCTGTGGACAGGGAGCCCTCCGGCTGGCAGTGGAGAGTAC ACAGCAGAGGAACGTGCATCAGGGCTGCACAGGCATGCCGTATCTCTCTTGTCAACACCTCTT GCCCTGTATGACGAGAACACCAAGCTCTGGTATGCTCAAATCGCACCATCACCCTGATGACAAGAT GTCCTCCGGCTCAACTACCGGATGAGGTTCTATTCACCAATTGGATGGAACCAACGACAATGAGC AGTCAGTGTGGCGTCACTCTCAAAGAACGAGAAAAATGGCTACGAGAAAAAAAGATTCCAGATGCA ACCCCTCTCCTGATGCCAGCTCACTGGAGTATCTGTTGCTCAGGGACAGTATGATTGGTAAATGC CTGGCTCTTATTGAGACCCCCAAGACCGAGCAGGATGGACATGATATTGAGAACGAGTGTCTAGGGAT GGCTGTCTGGCCATCTCACACTATGCCATGATGAGAACGATGAGTTGCCAGAACTGCCAAGGACA TCAGGTAAAGCGATATATTCCAGAACATTGATAAGTCATCAGACAGAGAACCTCTCACCAGGAT GCGGATAAATAATGTTTCAAGGATTCTAAAGGAATTAAACAACAAGACCAATTGACAGCAGCGT GTCACGCGATGACCTGAAGGTAAACTTGCTACCTTGGAAACTTGACAAAACATTACGGTGCTGA AATATTGAGACTTCATGTTACTGATTGATTCATCAGAAAATGAGATGAAATTGGTTATTGAAATGACGGT GGAAACGTTCTACTACGAAGTGTGGTACTGGAACTTGGAAATCCAGTGGAGGACAAACCAAA TGGTGTCTGTTGAAAAGGAAAAAAACTGAAGCGGAAAAACTGGAAAATAACACAAAGAAGGA TGAGGAGAAAAACAAAGATCCGGAGAGTGGAAACAATTCTTACTTCCCTGAAATCACTCACATTGT AATAAAGGAGTCTGTGGTCAGCATTAAACAAGCAGGACAACAAGAAAATGGAACGACTGAGCTCTTCCC CGAGGAGGCCCTGTCCTTGTGTCCTGGTAGATGGCTACTCCGGCTCACAGCAGATGCCATCATT ACCTCTGCACCGACGTGGCCCCCGTTGATCGTCCACAAACATACAGAATGGCTGTCATGGTCAATC TGACAGAATACGCCATCAATAAATTGCGGCAAGAAGGAAGCGAGGAGGGGATGTACGTGCTGAGGT GGGCTGCACCGACTTGACAAACATCTCATGACCGTCACCTGCTTGAGAAGTGTGAGCAGGTGAGG GTGCCAGAACGAGTTCAAGAACCTTCAGATCGAGGTGAGAAGGCCGCTACAGTGTGACGGTCC GGACCGCAGCTCCCCAGCTGGAGACCTCATGAGCCACCTCAAGAACGAGATCTGCGCACGGAT AACATCAGCTCATGCTAAACGCTGCTGCCAGCCAAAGCCCCGAGAAATCTCCAACCTGCTGGTGGC TACTAAGAAAGCCAGGAGTGGCAGCCGTCACCCATGAGCCAGCTGAGTTGATCGATGATCCTCA AGAAGGATCTGGTCAGGGCAGCACCTGGAGAGGCCAGGAGAACACACATCTATTCTGGACCC GATGGATTACAAGGATGACGAAGGAACCTCTGAAAGAGAACGATAAAAGTGTCTCAAAGTCTTAGA CCCCAGCCACAGGGATATTCCCTGGCTTCTCGAGGCAGCCAGCATGATGAGACAGGTCTCCACAA AACACATCGTGTACCTCATGGCTGTCGCCGACGTGGAGAATATCATGGTAAGAGATTGTTG GAAGGGGGTCTCTGGATCTTCTGACGCCAGTGCCTGAGCTACTTGGAGGATAAACGACTGGCCATGGAAATGTGT GTACTAAAACCTCCCTGGCCGTGAGGGCATGACAGTGTGAGCTGGCCGTTCATCAAGCTCAGT GACCCGGCATCCCATCGTGTCTAGGCAAGAACGATCTGAAACGAAATCCATGGATTGCTCC TGAGTGTGTTGAGGACTCCAAGAACCTGAGTGCTGCTGACAAGTGGAGCTTGGAAACACGCTC GGAAATCTGCTACAATGGCGAGATCCCTGAAAGACAAGACGCTGATTGAGAAAAGAGAGATTCTAT GAAAGCCGGTGCAGGCCAGTGACACCATCATGTAAGGAGCTGGCTGACCTCATGACCCGCTGATGA ACTATGACCCAATCAGAGGCCCTTCTCCGAGCCATCATGAGAGACATTAATAAGCTGAAAGAGCAGA ATCCAGATATTGTTCAAGAAAAAAACAGCAACTGAAGTGGACCCACACATTGAAAAGCGCTCC TAAAGAGGATCCGTGACTTGGAGAGGGCCTTGGAGGTTGAGCTGTCAGGTATGACCCGAG AGGGGACAATACAGGGAGCAGGGCTGTTAAATCTGTAAGCCTGAGAGTGGAGGTAACACATA GCTGATCTGAAAAAGGAATCGAGATCTTAAAGAACCTCATGAGAACATTGTAAGTACAAGGA ATCTGCACAGAACGAGGAAATGGTATTAGCTCATGGAATTCTGCCCTGGAAAGCCTTAAGG AATATCTCCAAAGAATAAGAACAAATAACCTCAAACAGCAGCTAAATGCGTTGAGATTGAA GGGGATGGACTATTGGCTCTCGGCAATACGTTACCGGGACTTGGCAGCAAGAACGACTGCTTGT AGAGTGAACACCAAGTAAAATTGGAGACTTGGTTAACCAAGCAATTGAAACCGATAAGGAGTATT ACACCGTCAAGGATGACCCGGACAGCCCTGTGTTGGTATGCTCCAGAATGTTAATGCAATCTAAAT TTTATATTGCCCTGACGCTGGTCTTGGAGTCACCTGCACTGAGCTGCTGACTTACTGTGATT TTCTAGTCCCAGGGCTTGTCTGAAATGATAGGCCAACCCATGCCAGATGACAGTCACAAGACT TGTGAATACGTTAAAGAAGGAAACGCTGCCGTGCCACCTAACTGTCAGATGAGGTTATCAACT TATGAGGAAATGCTGGAAATTCAACCATCCAATCGGACAAGCTTCAAGAACCTTATTGAAGGATT AGCACTTTAAATAAGAACGATGAAATAACATTAAATTCCACAGATTCAA

TABLE 20 : Amino Acid Sequence from Mouse

Sagres Tag No. S00039	Seq ID No. 202	MQYLNKEDCNAMAFCAKMRSFKKTEVKQVVPEPGVEVTYLLDREPLRLGSGEY TAEELCIRAAQECSISPLCHNLFALYDESTKLWYAPNRIITVDDKTSRLHYRMRFYF TNWHTNDNEQSVWRHSPKKQNGYEKRVPEATPLLDASSLEYLFAQGQYDLIK CLAPIRDPKTEQDGHDIENECLGMALISHYAMMKMQLPELPKDISYKRYIPETL NKSIRQRNLLTRMRINNVFKDFLKEFNNKTICDSSVHDLKVYLATLETSTLTKHG AEIFETSMILLISSENELSRCHSNSDGNVLYEVMVGTGNLGIQWRQKPNVVVEKEKN KLKRKKLEYNHKKDDERNKLREEWNNFSYFPEITHIVIKESVVSINKQDNKNMELK LSSREEALSFVSLVDGYFRLTADAHYLCTDVAPPLIVHNIQNGCHGPICTEYAINKL RQEGSEEGMYVLRWSCTDFDNILMTVTCFEKSEVLGGQKQFKNFQIEVQKGRYSL HGSMDFPSLRDLMNHLKKQILRTDNISFVLRKCCQPKPREISNLLVATKKAQEWAQ PVYSMSQLSFDRILKKDIQGEHLGRGTRTHIYSGTLLDYKDEEGIAEEKKIKVILKVL DPSHRDISLAFFEAASMMRQVSHKHIVYLYGVCVRDVENIMVEEFVEGGPLDFMH RKSDALTTPWKFKVAKQLASALSYLEDKDLVHGNVCTKNLLAREGIDSDIGPFIKL SDPGIPVSVLTRQECIERIPWIAPECVEDSKNLSVAADKWSFGTTLWEICYNGEPLK DKTLIEKERFYESRCRPVTPSCHELADLMTRCMNYDPNQRPFRAIMRDINKLEEQ NPDIVSEKQPTTEVDPTHFEKRFKIRDLGEGHFGKVELCRYDPEGDNTGEQVAV KSLKPESGGNHIAIDLKKEIEILRNLYHENIVKYKGICMEDGGNGIKLIMEFLPSGSLKE YLPKNKNKINLKQQLKYAIQICKGMDYLGSRQYVHRDLAARNVLVESEHQVKIGDF GLTKAIETDKEYYTVKDDRDSPFWYAPECCLIQCKFYIASDWSFGVTLHELLTYCD SDFSPMALFLKMICGPTHGQMTVTRLVKTKEGKRLPCPPNCPCDEVYQLMRKCWEF QPSNRRTFQNLIEGFEALLK
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TABLE 21 : Amino Acid Sequence from Human

<u>Sagres</u> <u>Tag No.</u> S00039	<u>Seq. ID</u> <u>No.</u> 203	MQYLNKEDCNAMAFCAKMRSSKKTEVNLEAPEPGVEVIFYLSDREPLRLGSGEYTA EELCIRAAQACRISPLCHNLFALYDENTKLWYAPNRTITVDDKMSLRLHYRMRFYFT NWHGTNDNEQSIVWRHSPKKQKNGYEKKIPDATPLLDASSLEYLFAQGQYDLVKC LAIRDPKTEQDGHDIENECLGMAVLAISHYAMMKMQLPELPKDISYKRYIPETLNK SIRQRNLLTRMRINNVFKDFLKEFNNKTICDSSVSTHDLKVVKYLATLETLTKHGAEIF ETSMILLISSENEMNFHSNDGGNVLYYEVMTGNLGIQWRHKPNVVSVKEKNKL KRKKLENKHKKDEEKNKIREEWNNSYFPEITHIVIKESVVSINKQDNKKMELKLSSH EEALSFVSLVDGYFRLTADAHYLCTDVAPPLIVHNIQNGCHGPICTEYAINKLRQEG SEEGMYVLRWSCTFDNILMTVTCFEKSEQVQGAQKQFKNFQIEVQKGGRYSLHGS DRSFPSLGDLMSHLKKQILRTDNISFMLKRCCQPKPREISNLLVATKKAQEWPVYP MSQLSFDRILKKDLVQGEHLGRGTRTHIYSGTLMDYKDDEGTSEEKKIKVILKVLDP HRDISLAFFEAASMMRQVSHKHIVLYGVCVRDVENIMVEEFVEGGPLDFMHRKS DVLTPPWKFVAKQLASALSYLEDKDLVHGNVCTKNLLLAREGIDSEC GFP I KLSDP GIPITVLSRQECIERIPWIAPECVEDSKNLSVAADKWSFGTTLWEICYNGEIPLKDKTLI EKERFYESRCRPVTPSCHELADLMTRCMNYDPNQRPFFRAIMRDINKLEEQNPDIVS EKKPATEVDPTHFEKRFKLKRIDLGEGHFGKVELCRYDPEGDNTGEQVAVKSLKPE SGGNHIADLKKEIEILRNLYHENIVKYKGICTEDGGNGIKLIMEFLPSGSLKEYLPKNK NKINLKQQQLKYAVQICKGMDYLGSRQYVHRDLAARNVLVESEHQVKIGDFGLTKAI TDKEYYTAKDDRDSPFWYAPECLMGSKFYIASDVWSFGVTLHELLTYCDSDSSPM ALFLKMI GPT HGQMTVTRLVNTLKEGKRLPCPPNC PDEVYQLMRKCWEFQPSNRT SFQNLIEGFEALLK
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Table 22 : Sagres Tag No. S00039 Nucleotide Sequence

<u>Sagres Tag No.</u>	<u>Seq ID No.</u>	Nucleotide Sequence
S00039	204	ACAAGACTTGAAGCGGTTCTGAAGAGGATTCGTACTGGAG AGGGTCACTTGGAAAGGTTGAGCTCTGCAGATATGATCCTGAGGG GACAACACAGGGAGCAGGTAGCTGTCAAGTCCCTGAAGCCTGAGA GTGGAGGTAACCACATAGCTGATCTGAAGAAGGAGATAGAGATCTTA CGGAACCTCTACCATGAGAACATTGTGAAGTACAAAGGAATCTGCAT GGAAGACGGAGGCAATGGTATCAAGCTCATCATGGAGTTCTGCCTT CGGGAAGCCTAAAGGAGTATCTGCCAAAGAATAAGAACAAAATCAAC CTCAAACAGCAGCTAAAAATGCCATCCAGAATTGTAAGGGGATGG ACTACTGGTTCTCGCAATAAGTTACCCGGACTTAGCAGCCAGA ATGTCCTTGTGAGAGTGAGCATCCAGTTGAGATTGGAGACCTTGGG TTAACCCAAGCCATTGAAACGATTAGGAGTACTACACAGTTCAGGAC CACCGGGAAAAGCCAGTGTCCGGTACGCTCCGGAATGTTAACCCA GTGTTAATTAAAACGCCCTCGATGTCCGGTCCTTGGAGTGACACT GCACGAGCTGCTCAATTACTGTGACTCCGAATTAGTCCCATTGGCCTT GGTCCCAGGTAAGCCAACTCCAGGCCAGAACAGAACATTGAAG GCCTGTGGATCACTGAAAGAAGGAAAGCCCTGGCATGTCCACCCAAAT GTCCTGATGAAGTTAACAGCCTATGGGAAAATTCCCTGGAATTGANCT ACTAACCGAACATTTCGGAACCTATGGAAGAGTTAACGCCCCTTA AATAGAAGCCTGGCACACTTAATCCCCATTCAAATCTTCTCCAAG CCTTAAAAAGGTTAACAGGAAAGTTGAATGGGCTAACGCCCCAAA AACCGCGGTACAATTGCAATTACGGGTCC

- 5 The Neurogranin nucleic acid and amino acid sequences of the invention are depicted in Tables 23, 24, 25, 26 and 27. The nucleic acid sequence shown in Table 23 is from mouse. The nucleic acid sequence shown in Table 24 is from human. The amino acid sequence shown in Table 25 is from mouse. The amino acid sequence shown in Table 26 is from human. The sequence of Sagres Tag No. S00092 is shown in Table 27.

TABLE 23 : Neurogranin Nucleic Acid Sequence from Mouse

<u>Sagres Tag No.</u>	<u>Seq. ID No.</u>	
S00092	205	GTTGGTCCTCGCTCCAGTTCTCCCCGCCACCCCTGCAGAAAGTGTCTCTGATTGGCT TCGAGGCCGCAGGGCTCAGGTTACATTGCAAGAGTTGCGGAGCGCGGGAGACCGG ACCCAAGAGGGAGAGAGGGCTGGTCTGCAAGGATTCTGCGCTGGTCGGGGAGTGC GACAGCCCCTGAGCTGCCACCCAGCATCGTACAAACCCACCCCCCTGCGCCAGG CTCCACCCCAGCCAAGGACCCCTAACACCGGAATGGACTGCTGCACGGAGAGCG CTGCTCCAAGCCAGACGACGATATTCTTGACATCCCGCTGGATGATCCGGAGCAA CGCCGCTGCAGCCAAAATCCAGGCAGTTCCGGGCCACATGGCGAGGAAGAAGA TAAAGAGCGGGAGAGTGTGGCCGGAAGGGACCGGGCCCCGGGGGACCGAGCGGAGC TGGGGGCGCCCCGGGGAGGCAGCGGGCGGGCCCCAGCGGAGACTAGGCCAGAGC TGAACGTTTAGAAGTCCAGAGGAGAGTGGATGCCCGTCCCCCTCGCAGTACA AGACTTCCCTACTGTGTTGTAGCCCCCTCTTCCCACCAACCAGCCAGCTTCAGGAG CCCCCCCCCTCCCCCGCCGCTCCAGAGACTCCCTCTCCCAAGGCTGGCTCGTCT TGGCGTAGCAAGTCCGTGCCCTTTAGCTCTCAGTCTAAC721GTGGCTCCTTT GCCTTTCTCCCACCCCTGTCCTAAACCCATACTCCAAAATGTCCTTTGCTTCAGCC CACCTGTCCACGCGCCAGCATGCAGCTGCTCCGAGCCTCGTGCCTCGCT GCGCGTACTGCAGAGGGCGCCAAATGCGTCGCCAAACTCTCAAAAAAAAGAAAGA AAAAAAAGAAAAAGAAAGAAAGAAAAAAAGCAACCACCAAGTCCTTCGTTCTGTGG GCAACGAAAGGGGGCGCCCGCTTTCCACCCTAGCCTAACCTAACCTCTAAAC CTGGGGCTAGGAAAGAGGGGAGGGAGGTTTCATGGTTATCTGATAATTCCCTTGCTC AAATGGAAAGTGAAGTCCTATCCCATACCTGCTGCCTCACCCCTTTTTCTGAAAACG CACCCCTGAGAGCAGCCCCCTCCCGCTCTTCTTGTATGCAAAGCCTCTGAGCGCC TGGAGGCTCCGGCAGGAGGAGACTCCGCAGCCCCGCCCATGATAGCCTCTCCCC CGTTGGGCTCTCGGGTTGTGGCTGGAAGGCTTTAATCTGCGTGTGCATGTTACC ATACTGGGTTGGAATGTGAATAAAAGAGGAATGTCGAAGTGT

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TABLE 24 : Neurogranin Nucleic Acid Sequence from Human

<u>Sagres Tag No.</u>	<u>Seq. ID No.</u>	GGCACGAGGCGCCAGCCTCGTCCCCGAGAGGACCCCCGACACCAGCATGGACT GCTGCACCGAGAACGCCCTGCTCCAAGCCGGACGACGACATTCTAGACATCCGCTGG ACGATCCCGCGCCAAACGCCGGCCGCCAAAATCCAGGCAGTTTCGGGGGCCAC ATGGCGCGGAAGAAGATAAAAGAGCGGAGAGCGCGGCCGAAAGGGCCGGCCCTG GGGGGCCTGGCGGAGCTGGGTGGCCCCGGGAGGCAGGGCGGGCGGGCCAGCG GAGACTAGGCCAGAAGAACTGAGCATTCAAAGTCCCGAGGAGAGATGGATGCCG CGTCCCCCTCGCAGCGACGAGACTCCCTGCCGTGGTGACCCCCCTCGCCAG CAACCTGCCAGCTACAGGAGCCCCCTGCCTCCAGAGACTCCCTCACCCAGGCAGG CTCCGTCCGGAGTCGCTGAGTCCGTGCCCTTAGTTAGTTCTGCAGTCTAGTATGG TCCCCATTGCCCTTCACTCCACCCACCTAAACCATGCGCTCCAACTTCCCTCT TTTGCTCTCGCCCACCTCTTCCCGACCCAGCATGCAGCTCTGCCCTCCGAGCCTCA GTGCGCTTCCGTCCGGCACTGCGGAGGGCGCCCTAACCGTCACCCAAAGCACACTCA CTTAAAGAAAAAACGAGTTCTTGTGCGCAGCTAAAGGGGCGCCCTACATC TCCGTGCCACTCCGCCCCAGCCTAGCCCCAAGACTTGGATCCGGGGAGATGAA GGGAAGAGGGTTGTTTGGTTTGGACGACCCCTGCTCTGACCGGAAGAGAAGTCCC TATCCCACACCTGCCGTACGTTCCCTCCCTTCCCCAGCGCACTGTTAGGGCAG CCTCTCCAGCTCTTGTGTTATGCAAACGCCGAGCGCCTGGGAGGCTCGGTAGGAGG AGTCTTCCACGGCCCCGCCCTGCGGTCCTCCCTCCCCCGCCGGGCT CCTGGGGCTGTGGCCGAAAGGTTCTGATCTCCGTGTGCATGTGACTGTGCTGGG TTGAATGTGAACAATAAGAGGAATGTCCAAGTGA
S00092	206	GGCACGAGGCGCCAGCCTCGTCCCCGAGAGGACCCCCGACACCAGCATGGACT GCTGCACCGAGAACGCCCTGCTCCAAGCCGGACGACGACATTCTAGACATCCGCTGG ACGATCCCGCGCCAAACGCCGGCCGCCAAAATCCAGGCAGTTTCGGGGGCCAC ATGGCGCGGAAGAAGATAAAAGAGCGGAGAGCGCGGCCGAAAGGGCCGGCCCTG GGGGGCCTGGCGGAGCTGGGTGGCCCCGGGAGGCAGGGCGGGCGGGCCAGCG GAGACTAGGCCAGAAGAACTGAGCATTCAAAGTCCCGAGGAGAGATGGATGCCG CGTCCCCCTCGCAGCGACGAGACTCCCTGCCGTGGTGACCCCCCTCGCCAG CAACCTGCCAGCTACAGGAGCCCCCTGCCTCCAGAGACTCCCTCACCCAGGCAGG CTCCGTCCGGAGTCGCTGAGTCCGTGCCCTTAGTTAGTTCTGCAGTCTAGTATGG TCCCCATTGCCCTTCACTCCACCCACCTAAACCATGCGCTCCAACTTCCCTCT TTTGCTCTCGCCCACCTCTTCCCGACCCAGCATGCAGCTCTGCCCTCCGAGCCTCA GTGCGCTTCCGTCCGGCACTGCGGAGGGCGCCCTAACCGTCACCCAAAGCACACTCA CTTAAAGAAAAAACGAGTTCTTGTGCGCAGCTAAAGGGGCGCCCTACATC TCCGTGCCACTCCGCCCCAGCCTAGCCCCAAGACTTGGATCCGGGGAGATGAA GGGAAGAGGGTTGTTTGGTTTGGACGACCCCTGCTCTGACCGGAAGAGAAGTCCC TATCCCACACCTGCCGTACGTTCCCTCCCTTCCCCAGCGCACTGTTAGGGCAG CCTCTCCAGCTCTTGTGTTATGCAAACGCCGAGCGCCTGGGAGGCTCGGTAGGAGG AGTCTTCCACGGCCCCGCCCTGCGGTCCTCCCTCCCCCGCCGGGCT CCTGGGGCTGTGGCCGAAAGGTTCTGATCTCCGTGTGCATGTGACTGTGCTGGG TTGAATGTGAACAATAAGAGGAATGTCCAAGTGA

TABLE 25 : Neurogranin Amino Acid Sequence from Mouse

<u>Sagres Tag No.</u> S00092	<u>Seq. ID No.</u> 207	MDCCTESACSKPDDILDIPPLDDPGANAAAQASFRGHMARKKIKSGECGRKGPGPGG PGGAGGGARGGAGGGPSGD
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TABLE 26 : Neurogranin Amino Acid Sequence from Human

5	<u>Sagres Tag No.</u> S00092	<u>Seq. ID No.</u> 208	MDCCTENACSKPDDILDIPPLDDPGANAAAQASFRGHMARKKIKSGERGRKGPGPGG PGGAGVARGGAGGGPSGD
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TABLE 27 : Sagres Tag No. S00092 Nucleic Acid Sequence

<u>Sagres Tag No.</u> S00092	<u>Seq. ID No.</u> 209	GTCAAAATACTGAGAATTAGAGGCATTGGATGCCAAGTCATAGAGAGGGACACATATA TACCAATACTTCCAAGGCTCAGGAAACATCATGGAAGAAGGGTAGGAAGAATTAAAN AACCGAGAAGAAGGGGGTGAGGTATGGAATGATGATTCCAGTCATGACTGGCTATT GAGTTAACACAGCTGGATCACCTGCACAAGATCTCCACAAGAGTGGGCCATTAAACA CTCTATCATGGAAAGAGGGAGGGCNTATGAGGTACCACCCCACCCCTGAAGATTATAC ACAATTAAATANTGGTGAGGTAGGGAGAGACATTACTTTAGGGTGCACTAGT ACAGTGCCTAC
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10 The Nrf2 nucleic acid sequences of the invention are depicted in Tables 28 through 31.

A Nrf2 nucleic acid sequence of the invention is depicted in Table 28 as SEQ ID NO. 210. The nucleic acid sequence shown is from mouse.

TABLE 28

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SEQ ID NO. 211 (in Table 29) represents the amino acid sequence of a protein encoded by SEQ ID NO. 210.

10

TABLE 29

MOUSE	
SEQ ID#	SEQUENCE
211	MDLIDILWRQDIDLGVSREVFDQSQRQKDYELEKQQKLEKERQEQLQKEQEKAFFFAQFQLDEETGEFLPIQPQAQHQIQTDTSGSASYSQVAHIPKQDALYFEDCMQLLAETFPFVDDHESLALDIPSHAESSVFTAPHQAQSLNSLEAAMTDLSSIEQDMEQVWQELFSIPELQCLNTENKQLADETTAVPSPEATLTEMDSNYHFYSSISSLEKEVGNCGPFHGLGFEDSFSSILSTDDASQLTSDLSNPTLNTDFGDEFYSAFIAEPSDDGSMPSSAISQSLSELLDGTIEGCDLSLCKAFNPKHAEGTMEFNDSDDGSLNTSPSRASPEHSVESSIYGDPPPGFSDSEMEELDSAPGSVKQNGPKAQPAHSPGDTVQPLSPAQGH SAPMRESQCENTTKKEVPVSPGHQKAPFTKDKHSSRLEAHLTRDELRAKALHIPFPVKEIIINLPVDDFNEMMSKEQFN EAQLALIRDIRRRGKNKAAQNCNRKRKLENIVELEQDLGHLKDEREKLLREKGENDRNLHLLKRRRLSTLYLEVFSMLR DEDGKPYSPSEYSLQ QTRDGNVFVLPKSKKPDTKNN

Table 30 (SEQ ID NO: 212) depicts a human Nrf2 nucleic acid sequence of the invention.

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TABLE 30

SEQ ID#	HUMAN
	SEQUENCE
212	TTGGAGCTGCCGCCGGGACTCCGCCCAGCAGGACATGGATTGACATACTTGGAGGAAGATAT AGATCTGGAGTAAGTCAGAAGTATTGACTTCAGTCAGCGACGGAAAGAGTATGAGCTGGAAAAACAGAAAAAA ACTTGAAAAGGAAAGACAAGAACAACTCCAAAAGGAGCAAGAGAAAGCCTTTCACTCAGTTACAACAGATGA AGAGACAGGTGAATTCTCCCAATTAGCCAGCCCAGCACACCAGTCAGAAACCAGTGGATCTGCCAACTACT CCCAGGGTGCACATTCCAAATCAGATGCTTGACTTGTACTGCAGCTTGGCAGACATTC CGTTTAGATGACAATGAGGTTCTCGGCTACGTTCTAGTCAGTTCTGATATTCCGGTCAACATCGAGA GCCAGTCTCATTGCTACTAATCAGGCTCAGTCACCTGAAACTCTGCTCAGGTAGCCCCGTTGATTAG ACGGTATGCAACAGGACATTGAGCAAGTTGGGAGGCTATTATCCATTCTGAGTTACAGTGTCTTAATATTG AAAATGACAAGCTGGTGAGACTACCATGGTCCAAGTCCAGAAGCCAAACTGACAGAAGTTGACAATTATCATT TTAATCTATACCCCTCAATGGAAAAAGAAGTAGGTAACGTAGTGTACCTTCTAATGCTTGGGATATTCC TTCAAGCAGCATCCTCTCACAGAACAGACCAACCCAAACAGTGCAGTCATTAATTACAGATGCCACAGTCAC ACAGATTGGTGAATTCTGCTTCTAGTCAGGCTCAGTCAGTGCATTAATTGGGGATATTCTGGGGATATGG CTTAAAGCATTCACTCTGAACCTCTAAATGGGCCATTGATGTTCTGATCTACACTTGAAGCTTCAA CCAAAACCACCCCTGAAAGCACAGCAGAAATTCAATGATTCTGACTCCGGCACTAAACACAAGTCCCAGTGT GGCATCACCAGAACACTCAGTGGATCTCCAGCTATGGAGACACACTACTTGGCCTCAGTGTGATTCTGAAGTGG AAGAGCTAGATAGTGGCCCTGGAAGTGTCAAACAGAAATGGCTCAAACACAGTACATTCTGGGGATATGG TACAACCCCTGTCACCTCTCAGGGCAGAGCAGTCACCTGCATGATGCCAATGTGAGAACACACAGAGAAA GAATTGGCTGTAAGTCTGGTATCGGAAAACCCATTCAACAAAGACAAACATTCAAGCCGCTGGAGGCTCAT CTCACAAAGAGATGAACCTAGGGCAAAGCTCTCCATATCCCATTCCCTGTAAGAAAAAAATCATTAACCTCCCTGTT GTTGACTTCAACGAAATGATGTCAAAGAGCAGTTCAATGAAGCTCAACTTGCTTAAATTGGGATATACTGAGG AGGGTAAGAATAAAAGTGGCTCAGAATTGAGAAAAGAAAATGGAAAATATAGAACTAGAGCAAGAT TTAGATCATTGAAAGATGAAAAGAAAATTGCTCAAAGAAAAGGAGAAAATGACAAAAGCCTTCACCTACTG AAAACAAACTCAGCACCTTATCTCGAAGTTTCACTGGCATGCTACGTGATGAAGATGGAAAACCTTCTCCTAG TGAATACTCCCTGCAAGAACAGAGATGGCAATGTTCTGCTGAGCTAGTTTTGTACTATTACTAAAGCTCCTACTGTGATG AAACTAGATTAGGAGGATTGACCTTCTGAGCTTCTGAGCTAGTTCTGAGCTAGTTTTGTACTATTACTAAAGCTCCTACTGTGATG TGAAATGCTCATACTTTATAAGAATTCTAGCAGGAAACTAGTATAGAAAATAACGAAACTTTA AAAAGCATGGAGTGTCACTGTTGAATCAGTAGTTCACTTAACTGAAACAAATTCTTAGGACACCATTTG GCTAGTTCTGTGTAAGTGTAAACTACAAAACATTATTATAGTGTCTTAGTGTATTAGATT ATATGATGATGACATCTGGCTAAAAGAAATTATTGCAAACACTAACCCAGATGACTTTTATAAAACTGTAT GGACAAAAATGGCTTTTATAATTAAATTGTTAGCTCTGGCAAAAAAAAAAATTNTTAAGAGCTGGTACTA ATAAAGGATTATTGACTGTTAAAAAAAAAAAAAA

Table 31 (SEQ ID NO: 213 depicts the amino acid sequence encoded by the nucleic acid sequence of SEQ ID NO: 212).

10

TABLE 31

SEQ ID#	HUMAN
	SEQUENCE
213	MDLIDILWRQDIDLGVREVDFDFQSRRKEYLEKOKKLEKERQEQLQKEQEKAFFTQLQLDEETGEFLPIQPAQHTQS ETSGSANYSQVAHIPKSDALYFDDCMQLLAQTFPVDDNEVSSATFQSLVPDPIGHIESPVFIATNQAQSPETSVAQVA PVLDGMQQDIEQWEEILLSIPELQCLNIENDLKVETTMVPSPEAKLTEVDNYHFYSSIPSMEKEVGNCSPHFLNAFE DSFSSILSTEDPNQLTVNSLNSDATVNTDFGDEFYSAFIAEPSISNSMPSPATLSHSLSELLNGPIDVSDLCKAFNQN HPESTAEOFNDSDSGISLNTPSPVASPEHSVESSSYGDTLLGLSDSEVEELDSAPGSVKQNGPKTPVHSSGDMVQPLS PSQQQSTHVDAQCENTPEKELPVSPGHRKPTFKDKHSSRLEAHLTRDELRAKALHIPFPVKEIINLPVVDFNEMMS KEQFNEAQLALIRDIRRGNKNVAAQNCRKRKLENIVELEQQLDHLKDEKEKLLKEKGENDKSLHLLKKQLSTLYEVF SMLRDEDGKPYPSEYSLQQTRDGNVFLVPKSKKPDVKNN

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All accession numbers cited herein are incorporated by reference in their entirety. All references cited herein are expressly incorporated in their entirety by reference.

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CLAIMS

We claim:

1. A recombinant nucleic acid comprising a nucleotide sequence selected from the group consisting of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212).
2. A host cell comprising the recombinant nucleic acid of claim 1.
3. An expression vector comprising the recombinant nucleic acid according to claim 2.
4. A host cell comprising the expression vector of claim 3.
5. A recombinant protein comprising an amino acid sequence selected from the group consisting of the sequences outlined in Table 14 (SEQ ID NO: 194), Table 5 (SEQ ID NO: 179), Table 7 (SEQ ID NO: 181), Table 9 (SEQ ID NO: 183), Table 10 (SEQ ID NO: 186), Table 11 (SEQ ID NO: 188), Table 12 (SEQ ID NO: 190), Table 13 (SEQ ID NO: 192), Table 16 (SEQ ID NO: 197), Table 17 (SEQ ID NO: 199), Table 20 (SEQ ID NO: 202), Table 21 (SEQ ID NO: 203), Table 25 (SEQ ID NO: 207), Table 26 (SEQ ID NO: 208), Table 29 (SEQ ID NO: 211), and Table 31 (SEQ ID NO: 213).
6. A method of screening drug candidates comprising:
 - a) providing a cell that expresses a lymphoma associated (LA) gene selected from the group consisting of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212), or fragment thereof;
 - b) adding a drug candidate to said cell; and
 - c) determining the effect of said drug candidate on the expression of said LA gene.
7. A method according to claim 6 wherein said determining comprises comparing the level of expression in the absence of said drug candidate to the level of expression in the presence of said drug candidate.
8. A method of screening for a bioactive agent capable of binding to an LA protein (LAP), wherein said LAP is encoded by a nucleic acid selected from the group consisting of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212), said method comprising:
 - a) combining said LAP and a candidate bioactive agent; and
 - b) determining the binding of said candidate agent to said LAP.
9. A method for screening for a bioactive agent capable of modulating the activity of an LA protein (LAP), wherein said LAP is encoded by a nucleic acid selected from the group consisting of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212), said method comprising:
 - a) combining said LAP and a candidate bioactive agent; and
 - b) determining the effect of said candidate agent on the bioactivity of said LAP.
10. A method of evaluating the effect of a candidate lymphoma drug comprising:
 - a) administering said drug to a patient;

- 5 b) removing a cell sample from said patient; and
c) determining alterations in the expression or activation of a gene selected from the group consisting of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212).
- 10 11. A method of diagnosing lymphoma comprising:
a) determining the expression of one or more genes selected from the group consisting of a nucleic acid of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212), or a polypeptide encoded thereby in a first tissue type of a first individual; and
b) comparing said expression of said gene(s) from a second normal tissue type from said first individual or a second unaffected individual;
wherein a difference in said expression indicates that the first individual has lymphoma.
- 20 12. A method for inhibiting the activity of an LA protein (LAP), wherein said LAP is encoded by a nucleic acid selected from the group consisting of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212), said method comprising binding an inhibitor to said LAP.
- 30 13. A method of treating lymphoma comprising administering to a patient an inhibitor of an LA protein (LAP), wherein said LAP is encoded by a nucleic acid selected from the group consisting of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212).
- 40 14. A method of neutralizing the effect of an LA protein (LAP), wherein said LAP is encoded by a nucleic acid selected from the group consisting of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212), comprising contacting an agent specific for said LAP protein with said LAP protein in an amount sufficient to effect neutralization.
- 45 15. A polypeptide which specifically binds to a protein encoded by a nucleic acid of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212).
- 50 16. A polypeptide according to claim 15 comprising an antibody which specifically binds to a protein encoded by a nucleic acid of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8 (SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212).
- 56 17. A biochip comprising one or more nucleic acid segments selected from the group consisting of a nucleic acid of the sequences outlined in Tables 14 (SEQ ID NO: 193), 4 (SEQ ID NO: 178), 6 (SEQ ID NO: 180), 8

(SEQ ID NO: 182), 9 (SEQ ID NO: 183), 10 (SEQ ID NO: 185), 11 (SEQ ID NO: 187), 12 (SEQ ID NO: 189), 13 (SEQ ID NO: 191), 15 (SEQ ID NO: 195), 16 (SEQ ID NO: 196), 17 (SEQ ID NO: 198), 18 (SEQ ID NO: 200), 19 (SEQ ID NO: 201), 22 (SEQ ID NO: 204), 23 (SEQ ID NO: 205), 24 (SEQ ID NO: 206), 27 (SEQ ID NO: 209), 28 (SEQ ID NO: 210) and 30 (SEQ ID NO: 212).

- 5 18. A method of diagnosing lymphomas or a propensity to lymphomas by sequencing at least one LA gene of an individual.
19. A method of determining LA gene copy number comprising adding an LA gene probe to a sample of genomic DNA from an individual under conditions suitable for hybridization.

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization
International Bureau



(43) International Publication Date
28 March 2002 (28.03.2002)

PCT

(10) International Publication Number
WO 02/024867 A3

(51) International Patent Classification⁷: **C12Q 1/68**, C07K 14/47, C12N 5/10, 15/85

(21) International Application Number: PCT/US01/29798

(22) International Filing Date:
24 September 2001 (24.09.2001)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:
09/668,644 22 September 2000 (22.09.2000) US
09/905,390 13 July 2001 (13.07.2001) US
09/905,491 13 July 2001 (13.07.2001) US

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(81) Designated States (*national*): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.

(84) Designated States (*regional*): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

— with international search report

(88) Date of publication of the international search report:
14 August 2003

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.



WO 02/024867 A3

(54) Title: NOVEL COMPOSITIONS AND METHODS FOR LYMPHOMA AND LEUKEMIA

(57) Abstract: The present invention relates to novel sequences for use in diagnosis and treatment of lymphoma and leukemia. In addition, the present invention describes the use of novel compositions for use in screening methods.

INTERNATIONAL SEARCH REPORT

In Application No
PCT/US 01/29798

A. CLASSIFICATION OF SUBJECT MATTER
IPC 7 C12Q1/68 C07K14/47 C12N5/10 C12N15/85

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)
IPC 7 C07K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

BIOSIS, EMBASE, EPO-Internal, WPI Data, PAJ, EMBL

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>LI JIAYIN ET AL: "Leukaemia disease genes: Large-scale cloning and pathway predictions." NATURE GENETICS, vol. 23, no. 3, November 1999 (1999-11), pages 348-353, XP002225264 ISSN: 1061-4036 the whole document</p> <p>---</p> <p>-/-</p>	1-19

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- "&" document member of the same patent family

Date of the actual completion of the international search 17 December 2002	Date of mailing of the international search report 25.04.2003
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl Fax: (+31-70) 340-3016	Authorized officer Schalich, J

INTERNATIONAL SEARCH REPORT

Inter Application No
PCT/US 01/29798

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	SORENSEN ANNETTE BALLE ET AL: "Sint1, a common integration site in SL3-3-induced T-cell lymphomas, harbors a putative proto-oncogene with homology to the septin gene family." JOURNAL OF VIROLOGY, vol. 74, no. 5, March 2000 (2000-03), pages 2161-2168, XP002225265 ISSN: 0022-538X the whole document ---	1-19
X	HALLEK M ET AL: "REDUCED RESPONSIVENESS OF ADENYLYLATE CYCLASE TO FORSKOLIN IN HUMAN LYMPHOMA CELLS" BIOCHEMICAL PHARMACOLOGY, vol. 42, no. 7, 1991, pages 1329-1334, XP008011750 ISSN: 0006-2952 the whole document	1-19
Y	---	10,11, 13,18
X	DATABASE EMBL [Online] SQ 23, 28 October 1999 (1999-10-28) retrieved from EMBL Database accession no. AAZ41972 XP002225267 abstract; claim 3	1-9,12, 14,20-22
X	& DE 198 17 947 A (METAGEN GES FUER GENOMFORSCHUNG) 28 October 1999 (1999-10-28)	1-9,12, 14,20-22
Y	the whole document	10,11, 13,18
X	---	1,5,15
X	DATABASE EMBL [Online] 21 March 1996 (1996-03-21) retrieved from EMBL Database accession no. AAR94559 XP002225268 abstract; claim 2	1,5,15
X	& WO 96 08260 A (MOUNT SINAI SCHOOL MEDICINE) 21 March 1996 (1996-03-21)	1,5,15
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X	WO 98 16557 A (GEN HOSPITAL CORP) 23 April 1998 (1998-04-23)	15
X	the whole document	
X	---	
X	WO 94 02636 A (HITACHI CHEMICAL CO LTD ;MITSUHASHI MASATO (US); COOPER ALLAN (US)) 3 February 1994 (1994-02-03)	17,19
X	the whole document	

INTERNATIONAL SEARCH REPORT

ional application No.
PCT/US 01/29798

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:

2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:

3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

see additional sheet

1. As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.

2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.

3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:

4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

1-19 all partially

Remark on Protest

- The additional search fees were accompanied by the applicant's protest.
- No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: claims 1-19, partially

LA gene (SEQ.ID 193) and protein (SEQ.ID 194) and their use

2. Claims: claims 1-19, partially

LA gene (SEQ.ID 178) and protein (SEQ.ID 179) and their use

3. Claims: claims 1-19, partially

LA gene (SEQ.ID 180) and protein (SEQ.ID 181) and their use

4. Claims: claims 1-19, partially

LA gene (SEQ.ID 182)

5. Claims: claims 1-19, partially

LA gene (SEQ.ID 183) and protein (SEQ.ID 184) and their use

6. Claims: claims 1-19, partially

LA gene (SEQ.ID 185) and protein (SEQ.ID 186) and their use

7. Claims: claims 1-19, partially

LA gene (SEQ.ID 187) and protein (SEQ.ID 188) and their use

8. Claims: claims 1-19, partially

LA gene (SEQ.ID 189) and protein (SEQ.ID 190) and their use

9. Claims: claims 1-19, partially

LA gene (SEQ.ID 191) and protein (SEQ.ID 192) and their use

10. Claims: claims 1-19, partially

LA gene (SEQ.ID 195) its use

11. Claims: claims 1-19, partially

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

LA gene (SEQ.ID 196) and protein (SEQ.ID 197) and their use

12. Claims: claims 1-19, partially

LA gene (SEQ.ID 198) and protein (SEQ.ID 199) and their use

13. Claims: claims 1-19, partially

LA gene (SEQ.ID 200) and its use

14. Claims: claims 1-19, partially

LA gene (SEQ.ID 201) and protein (SEQ.ID 202) and their use

15. Claims: claims 1-19, partially

LA protein (SEQ.ID 203) and their use

16. Claims: claims 1-19, partially

LA gene (SEQ.ID 204) and its use

17. Claims: claims 1-19, partially

LA protein (SEQ.ID 205) and their use

18. Claims: claims 1-19, partially

LA gene (SEQ.ID 206) and protein (SEQ.ID 207) and their use

19. Claims: claims 1-19, partially

LA protein (SEQ.ID 208) and their use

20. Claims: claims 1-19, partially

LA gene (SEQ.ID 209) and its use

21. Claims: claims 1-19, partially

LA gene (SEQ.ID 210) and protein (SEQ.ID 211) and their use

22. Claims: claims 1-19, partially

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

LA gene (SEQ.ID 212) and protein (SEQ.ID 213) and their use

INTERNATIONAL SEARCH REPORT

Information on patent family members

Int	Application No
Fu, us	01/29798

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